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(54) Title: GENETIC ANALYSIS OF PEYER'S PATCHES AND M CELLS AND METHODS AND COMPOSITIONS TARGETING PEYER'S PATCHES AND M CELL RECEPTORS

(57) Abstract: Methods of increasing or decreasing the levels of a protein in a PP cell; methods of increasing antigen, vaccine, DNA vaccine delivery to M cells, use of human serum albumin and other transport enhancing proteins to enhance oral drug delivery; use of calreticulin to enhance oral antigen delivery, use of other cell surface proteins, receptors, and transporters to enhance delivery to M cells of antigens or vaccine delivery vehicles, use of other cytoplasmic proteins to regulate intracellular trafficking and delivery to mucosal immune sampling and processing systems.

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TITLE OF THE INVENTION

5 GENETIC ANALYSIS OF PEYER'S PATCHES AND M CELLS AND METHODS AND
COMPOSITIONS TARGETING PEYER'S PATCHES AND M CELL RECEPTORS

CROSS REFERENCE TO RELATED APPLICATIONS

10 This application claims the benefit of U.S. provisional application 60/281,387 filed April 4,
2001, and U.S. provisional application 60/302,591 filed July 2, 2001.

FIELD OF THE INVENTION

15 This invention relates to the genetic analysis of M cells and methods and
compositions targeting M cell receptors.

BACKGROUND OF THE INVENTION

20 The Peyer's patch of the intestinal lining is a specialized tissue that allows the
immune system to identify foreign antigens that require an immune response. It is also a
potential pathway for orally delivered drugs to cross the intestinal barrier into the
bloodstream. Central to these properties are M cells, which populate the patch's epithelial
sheet. In view of the importance of the Peyer's patch and its M cells for the immune
response and drug delivery, it is desirable to identify the cell proteins important for these
phenomena. It is also desirable to increase the amounts of such important proteins in order
25 to either facilitate the immune response and drug delivery or promote the conversion of non-
M cells to M cells.

Similarly, it is important to identify and further decrease the levels of proteins whose
absence or down-regulation in expression facilitates the immune response and drug delivery,
or promotes the conversion of non-M cells to M cells.

30

BRIEF SUMMARY OF THE INVENTION

Increasing the levels of a protein or antigen-protein combination

In a first general aspect, the invention is a method of increasing the levels of a protein

in a Peyer's patch cell, said method comprising delivering to said cell a nucleic acid coding for a protein, wherein absent said increase, the levels of said protein or its mRNA is greater than in a non-Peyer's patch cell.

5 Peyer's patch cells of particular interest are M cells. The levels of a protein or its mRNA in Caco-2 cells co-cultured with Raji B cells are considered herein to be representative of such levels in a human Peyer's patch M cell. Monoculture Caco-2 cells are considered herein to be an appropriate non-Peyer's patch cell for purposes of comparison of such protein or mRNA levels.

10 The levels of a protein or its mRNA in rat Peyer's patch epithelial cells can be compared to their respective levels in a culture of rat normal gut epithelial cells. Absent evidence to the contrary, results of rat cells are assumed to be predictive of the results in human cells.

15 The presence of Increased levels of an mRNA, and therefore presumptively its protein, are indicated in the Table 2 and 3 by a **, a *, or an expression Fold Change greater than 1.00. Preferred are those indicated by a ** or an expression Fold Change greater than 2.00. Most highly preferred are those indicated by a **. The presence of decreased levels of an mRNA, and presumptively its protein, are indicated by a minus sign (-) or an expression Fold Change less than 1.00. Preferred targets are those indicated by a minus sign or an expression Fold Change less than 0.50.

20 In embodiments of particular interest, the protein is a receptor, a transporter, cell surface antigen, or cell adhesion molecule, especially a receptor. In other embodiments of particular interest, the protein is selected from the group consisting of nucleoside diphosphate kinases and member of the 14-3-3 family.

25 In the methods of greatest interest, the nucleic acid is delivered to a human cell. There are many delivery options, one of which is to deliver it by the oral route with the cell in a human, another to deliver it to a cell outside a human.

30 In an important variation of the method, a nucleic acid coding for a tumor antigen or foreign peptide is also delivered to the Peyer's patch cell. The purpose of this aspect of the invention is to improve the immune response to a tumor antigen or the foreign peptide. Normally, therefore, the foreign peptide will be that of a virus or infectious microorganism. A tumor antigen is one that is more abundant in a tumor cell than its normal counterpart.

Decreasing the levels of a protein

Another general aspect of the invention is a method of decreasing the levels of a

protein in a Peyer's patch cell, said method comprising delivering to said cell an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference (RNAi) nucleic acid molecule, said anti-sense, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for said protein, wherein absent said anti-sense nucleic acid molecule, ribozyme or RNAi nucleic acid, the levels of said protein or its mRNA are less than in a non-Peyer's patch cell. More preferably the anti-sense nucleic acid is complementary to a sequence of at least 15 nucleotides of the mRNA of the protein, and most preferably to a sequence of at least 30 nucleotides of the mRNA of the protein. It is preferred that the protein is coded for by a gene with an expression Fold Change denoted by a minus sign (-) or an expression Fold Change less than 0.50.

In a particular embodiment, the latter method comprises delivering to said cell an anti-sense nucleic acid molecules, a ribozyme or RNAi nucleic acid molecules, said anti-sense, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for at least 5 different proteins, wherein absent said anti-sense, ribozyme or RNAi nucleic acid molecule, the levels of each of said proteins or its mRNA are less than in a non-Peyer's patch cell.

Alternatively described, the latter invention is a method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell an anti-sense nucleic acid molecule, ribozyme or RNAi nucleic acid molecules, said anti-sense, ribozyme or RNAi nucleic acid forming a double-stranded molecule with part or all of the mRNA for said protein, wherein absent said anti-sense, ribozyme or RNAi nucleic acid molecule, the levels of said protein or its mRNA are less than in a non-Peyer's patch cell.

Cells of the invention

A human or rat cell to which any of the above methods in this Brief Summary of the Invention section has been applied, or the progeny of said cell, is also an aspect of the present invention.

Delivery enhancement using a targeting ligand which targets a receptor, a transporter or a cell-surface molecule expressed on surface of M cells or Peyer's patch tissue cells

In another general aspect, the invention is a method of targeting an antigen or a drug delivery vehicle containing an antigen, or a drug delivery vehicle containing an antigen and adjuvant, or a drug delivery vehicle containing a drug, or a viral vector, or a bacterio-phage vector such as, but without limitation M13 or Fd, or a bacterial vector or a gene delivery

vector expressing an antigen of interest, or a viral vector, or a bacterio-phage vector such as, but without limitation M13 or Fd, or a bacterial vector or a gene delivery vector expressing a gene product(s) to M cells of Peyer's patch tissue, by targeted delivery to receptors, or to transporters or to other cell surface proteins which are found to be expressed on the cell surface of M cells or other cells found within Peyer's patch tissue, or which are found to be differentially expressed on these cells. Said gene product(s) coded by the viral vector, or a bacterio-phage vector such as, but without limitation M13 or Fd, or a bacterial vector or a gene delivery vector regulate the function of Peyer's patch cells to M cell phenotype or regulate M cell function to increase their immuno-surveillance or antigen presentation to the mucosal immune system.

In one embodiment, a phage display library such as M13 or Fd which express random peptide sequences on the surface of the phage, coded by example gene III or gene VII of M13 or Fd bacteriophage, can be screened by in vivo panning against example Peyer's patch tissue found in vivo in the GIT, in order to discover and identify phage or targeting ligands which specifically target M cells or Peyer's patch tissue in vivo in the GIT; such phage which target M cells and Peyer's patch tissue can subsequently be genetically engineered to encode a gene or genes of interest such as a DNA vaccine gene, a gene coding for an antigen of interest together with gene(s) which modify M cell function and which enhance the immuno-responsiveness of the M cells to the antigen or DNA vaccine product coded by the genetically engineered bacteriophage genome.

Delivery enhancement using transport enhancing proteins

Another invention disclosed herein is a method for enhancing transport of a drug through the gastrointestinal tract, said method comprising orally administering said drug in a composition that comprises a transport-enhancing protein, said transport-enhancing protein selected from the group consisting of human serum albumin (HSA), clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, and Ca^{2+} -dependant phospholipase A_2 (Ca^{2+} -pla2), or a homolog that has at least 80% amino acid identity with said transport-enhancing protein over a length of said transport-enhancing protein identical to the homolog. In a preferred embodiment, the homolog has at least 90% amino acid identity with the transport-enhancing protein over a length of the transport-enhancing protein identical to the homolog. In a more preferred embodiment, the transport-enhancing protein is selected from the group consisting of human serum albumin (HSA), clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, and Ca^{2+} -pla2.

Method of delivering a vaccine to a target cell

Further invention disclosed herein is a method of delivering a vaccine to a target cell, said method comprising utilizing as the target cell a Peyer's patch cell in which a protein or mRNA is upregulated.

5

Method of decreasing the levels of a protein

Yet, another invention disclosed herein is a method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell a DNA molecule coding for an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule (RNAi), said anti-sense molecule, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for said protein, wherein absent said anti-sense molecule, ribozyme or RNAi nucleic acid, the levels of said protein or its mRNA is less than in a non-Peyer's patch cell.

15

Method of increasing the extent to which the function of a protein is carried out

Another invention disclosed herein is a method of increasing the extent to which the function of a protein is carried out in a Peyer's patch cell, said method comprising delivering to said cell a nucleic acid coding for said protein, wherein absent said delivery, the level of said protein or its mRNA is greater in said cell than in a non-Peyer's patch cell.

20

Chimeric protein that comprises two or more segments, each of said segments enhancing a different step in the peptide transport process

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Another invention disclosed herein is a chimeric protein that comprises two or more segments, each of said segments enhancing a different step in the peptide transport process, said steps selected from the group consisting of binding to a cell such as an M cell, transporting the peptide into the cell such as an M cell, presenting the chimeric protein to a protein processing pathway within a cell such as an M cell in order to maximise processing in a way to optimize presentation of the processed chimeric peptides to epitopes suitable for immune activation, transporting the peptide through the cell such as an M cell, and transporting the peptide out of the cell such as an M cell to an underlying immune cell such as a B-cell or T-cell.

30

Delivery enhancement using calreticulin and other proteins

Another method disclosed herein is a method to facilitate intracellular trafficking of an antigen that has been orally delivered by itself or as part of a composition or particle, said method comprising administering calreticulin.

Related to the latter invention is a chimeric protein comprising the amino acid sequences for (1) calreticulin, rab family proteins and and/or a ribosomal protein, and (2) a second polypeptide. Also related is a method of administering a polypeptide, where said polypeptide is part of the chimeric protein and wherein said chimeric protein is orally administered.

DETAILED DESCRIPTION OF THE INVENTION

The present invention and the related research were intended to improve targeted vaccine delivery and targeted gene delivery methods, especially as they relate to Peyer's patch cells. In significant part, this was achieved by identifying proteins whose up-regulation or down-regulation would indicate their possible or probable role in cellular functions important to vaccine and or drug delivery. In some cases, such as receptors, the proteins are important from the point of view of cell specificity during the delivery process. In many cases, the proteins have functions that are important after the vaccine or drug enter the cell.

Closely related to those inventions and research goals, was the concept that in M cells there would be proteins that, as compared to M cell precursors, were up-regulated or down-regulated. The identification of such proteins provides a strategy for altering M cell precursors so as to shift their phenotype toward that of M cells.

As indicated, one aim of the research related to the present invention was to determine if there were detectable differences in protein/gene expression between: (1) Peyer's patch (PP) and non-Peyer's patch (NPP) rat gastrointestinal tract (GIT) tissue and (2) M cell enriched follicle-associated epithelium of Peyer's patch (PP FAE) tissue. This was done with a view to finding novel or highly expressed ligand targeting sites on the Peyer's patch or M cells as well as other protein relevant to the delivery of drugs across the GIT.

This invention is based in part on the discovery of over-expression of a range of genes in Peyer's patch (PP) tissue from rat small intestine in comparison to normal non-Peyer's patch (NPP) small intestine tissue.

This invention is also based on the discovery of over-expressed genes in co-cultures of Caco-2 cells. The idea was to use genetic mapping of the M cell co-culture, e.g. Caco-2

cells co-cultured with Raji cells versus a monolayer of Caco-2 cells, to ascertain the differences in epithelial gene expression between M cells and enterocytes. It became immediately apparent that some of these gene products are going to be unique apical membrane proteins (e.g. receptors, transporters, adhesion proteins) in M cells. By examining the differences between M cells and enterocytes in vitro and in vivo, one could identify key targets that can be used to generate M cell specific ligands. These ligands can then be used for targeting oral vaccines in particles.

The identification of over-expressed ribosomal proteins or homologues/related proteins thereof indicates a generally higher protein turnover or protein synthesis capacity in PPs or a possible role for such ribosomal proteins (or homologues thereof) in other cellular functions such as protein chaperoning, endocytosis, trafficking of proteins/antigens/particulates/viruses uptaken from the lumen of the gastrointestinal tract (GIT) and/or from the M cells to underlying immune cells, antigen presenting cells, dendritic cells, B cells, other cell types.

The identification of a series of transcription factors (TFs) that are over-expressed in PP tissue versus the control enterocyte GIT tissue is considered herein to indicate a role for such TFs in the development of M cell phenotype, in conferring M cell phenotype and/or in programming M cells to prime other downstream cellular events leading to a better or more efficacious immune outcome following antigen presentation. The co-delivery of genes coding for such TFs with either antigens themselves and /or with gene(s) coding for antigen(s) of question to M cells and/or PP tissue following oral administration provides the basis for a more efficacious and pronounced immune outcome when the TF coding genes are key or vital for driving M cells / PP tissue to an effective immune outcome.

The general over-expression of a number of proteins species in PPs versus NPPs, both membrane and cytosolic-associated was also determined by a novel technique of enrichment and M cell selection following enrichment of the follicle-associated epithelium (FAE) of Peyer's patch (PP FAE) by ethylene-diamine tetra-acetic acid (EDTA) extraction and recovery of M cells / PP FAE. Such novel or differentially expressed proteins have significant implications for the use of this protein expression information and methods of selection / enrichment of M cells / PP tissue for the targeting of drug/vaccine uptake to Peyer's patches. Among proteins found to be over expressed in rat PP tissue following this enrichment technique was the human serum albumin homolog which is considered here to have implications for drug / cargo transport from the GIT either into or across intestinal tissue including PP tissue and systemic delivery of same to the blood.

Incorporation by reference

All references cited herein are incorporated herein by reference in their entireties.

5 All GenBank records specified by their accession numbers are incorporated herein by their entireties.

The GenBank amino acid sequences and nucleotide sequences specified by their GenBank ID number are incorporated by reference herein. All GenBank records corresponding to those ID numbers are incorporated herein in their entirety. Absent a date specifying the date of the record, the date of the record is the filing date of this application.

10 Many of GenBank sequences specified by their GenBank ID numbers are reproduced herein in the section "Amino acid sequences and nucleotide sequences corresponding to selected GenBank ID numbers." The CDS line refers to the exon(s).

15 Any GenBank ID numbers specified herein, absent a decimal point and an integer following that decimal point, is for GenBank version 1 of that sequence. Any GenBank ID number that has a decimal point and an integer following it is the GenBank version number.

20 The invention will be illustrated in more details with reference to the following Examples, but it should be understood that the present invention is not deemed to be limited thereto.

EXAMPLESExample 15 Preparation of cytosolic (S100) and membrane (P100) proteins from rat PP and NPP tissues

Protein samples were prepared from PP and NPP tissue extracted from freshly sacrificed rats. These protein samples underwent electrophoresis on denatured SDS-PAGE gels and were stained using two different standard proteins Commassie Blue stains. Subsequently, fresh PP and NPP tissue samples were fractionated into cytosolic (S100) and
10 membrane (P100) proteins and these samples were also electrophoresed on SDS-PAGE in order to compare S100 and P100 fractions in both PP and NPP tissues.

Example 215 Preparation of GIT tissue or co-culture cell membrane (P100) and cytosolic (S100) fractions

The fractions were prepared using the following procedure:

1. Scrape the co-culture cells into PBS and pool cells into a universal.
2. Centrifuge the cells for 5 minutes at 1,500 rpm.
3. Remove the supernatant.
- 20 4. Re-suspend the cell pellet in 3 volumes of ice-cold HED buffer, and allow it to swell for 5 minutes on ice.
5. Homogenize the cells for 30 seconds.
6. Centrifuge the homogenate in hard walled tubes at 40,000rpm for 45minutes at 4°C in a Beckmann Ultra Centrifuge (rotor Ti90).
- 25 7. Remove the supernatant (S100) and re-suspend the pellet (P100) in 3 volumes of HEDG buffer, before centrifugation again at 1000rpm for 2min. Remove the supernatant and store on ice. Repeat the procedure and add the second supernatant to the first.
8. Determine the protein concentration (using the Bio-Rad protein assay).
9. All fractions were stored at -80°C.

30 The following reagents were used in the above methods:

HED buffer (20mM HEPES pH 7.67), 1mM EGTA, 0.5mM dithiothreitol, 1mM phenylmethylsulphonyl fluoride (PMSF):

HEPES (pH to 7.67)	0.5206g
EGTA	38.04mg
35 Dithiothreitol	7.71 mg

Distilled water to 100ml

10µl PMSF stock solution was added to 1ml of buffer prior to use.

HEDG buffer (the same as HED buffer plus 100mM NaCl, 10% glycerol)

5	NaCl	0.584g
	Glycerol	11.4ml
	HEPES (pH to 7.67)	0.5206g
	EGTA	38.04mg
	Dithiothreitol	7.71 mg

10	Distilled water	to 100ml
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10µl PMSF stock solution was added to 1ml of buffer prior to use.

PMSF(100mM) stock solution

	PMSF	17.42mg
15	Isopropanol	

Example 3

Isolation of epithelial sheaths from rat Peyer's patch and non-Peyer's patch tissue

20 The M cell is a very elusive cell type, at least in terms of isolating a purified population. Previous attempts have found that when M cells are separated and purified and put into culture they very quickly lose their characteristic morphology and probably gene/protein expression profile. In many cases this is due to the length of time taken to isolate and purify the cells from the very homogenous mix of cells in a Peyer's patch. We

25 desired a quick and routine method to enrich for M cells in Peyer's patch samples. M cells are only contained in the epithelium of Peyer's patches, the so-called follicle associated epithelium (FAE), while underneath the epithelial layer lays all the B and T lymphocytes, dendritic cells etc. So by isolating the epithelium away from the rest of the Peyer's patch dome, we are greatly enriching it for the M cell population. Previously, treatment of mouse

30 intestinal tissue with EDTA was shown to cause separation of the epithelium as a sheet from the rest of the tissue, allowing for its specific isolation (Bjerknes M and Cheng H (1981). Methods for the isolation of intact epithelium from the mouse intestine. *Anat. Rec.*, (199):565). This method was adapted for the isolation of FAE from rat Peyer's patch. Control epithelium from normal gut tissue (no Peyer's patches) was used as a control.

Epithelial sheaths were prepared using EDTA method comprising the following steps:

1. Sacrifice the rats (Wistar) by cervical dislocation.
2. Remove the entire length of the GIT tract from (but not including) the stomach to the caecum and place in a dish of PBS (at room temperature).
3. Excise the Peyer's patches, taking care to remove as much normal non-PP GI tissue as is visible. Rinse briefly in PBS.
4. Also take samples of normal non-Peyer's (NPP) tissue close to the patches, rinse in PBS and treat as for the PPs (steps 5-9, 11-12).
5. Pool the PP's from the entire GI section in Hank's Buffered Saline Solution (HBSS, Gibco Life Sciences) with 0.011M glucose and 25mM Hepes.
6. When pooling is complete, place PP sections into 15-20ml of HBSS (with 0.011M glucose and 25mM Hepes) along with 40mM of EDTA into a small conical flask.
7. Add a stirrer bar to the flask, place on a stirring plate and spin the PP's for 15 min at RT.
8. After 15 minutes pipette the PP solution vigorously with a wide-bore 3ml plastic pasteur pipette.
9. Strain the supernatant through a 100micron nylon cell strainer (from FALCON™, 352360).
10. Move the filter to another 50ml tube and wash out the residue material on the filter with HBSS (with 0.011M glucose and 25mM Hepes, no EDTA). This residue contains the majority of the PP dome epithelial sheaths.
11. Centrifuge the PP residue material at 3000 rpm for 5min. Also centrifuge the NPP tissue supernatant at 3000 rpm for 5min.
12. Snap freeze the cell pellets and store at -70°C.

Example 4

Identification of over-expressed proteins in enriched M cells / PP FAE cells

Epithelial cell layers of Wistar rat PP (representing enriched M cells / PP FAE cells) and normal villi were extracted using EDTA as described in Example 3 above. Epithelial layers from numerous Patches and rats were pooled and the protein isolated into either cytosolic and membrane fractions following centrifugal separation. 2D gel electrophoresis (between isoelectric points pH 3.5 to 10) was performed on 50 µg of each fraction, wherein the gels were silver stained. The gels were overlaid, and numerous differentially expressed

proteins between the membrane fractions of PP and normal villi epithelia were observed. Further, protein samples underwent a second 2D gel electrophoresis, this time the gel was stained with a "special" silver stain, that did not inhibit the mass spectrometry analysis of individual spots. The differentially expressed proteins were identified and highlighted by gel overlay.

Thirty-seven protein spots were identified that were increased in PP over villi epithelial membrane fractions. Of these, 16 spots (the most highly over-expressed) were chosen for mass spectrometry analysis. The spots were digested with endoproteinase Lys-C/trypsin (8:1 ratio) and analyzed on a MALDI-MS. The spots, however, gave very poor spectra and only 4 of the 16 were identifiable. These were:

- serum albumin;
- calreticulin;
- 14-3-3 zeta (tentative ID; mouse); and
- nucleoside diphosphate kinase B.

One protein showed homology to human serum albumin (HSA). Work by A. Fasano at Maryland had suggested that Zonulin, the human homologue of ZOT, showed sequence homology to human serum albumin (85% homology across the limited sequence available from the Fasano's work). Given our finding that a protein differentially expressed in rat PP tissue shows homology to HSA, we propose that HSA (or a homologue or splice variant thereof) is involved in drug transport in the GIT, in particular Peyer's patch tissue of the GIT.

Calreticulin is a 46-kDa Ca (2+)-binding chaperone of the endoplasmic reticulum membranes. This protein binds Ca (2+) with high capacity, affects intracellular Ca (2+) homeostasis, and functions as a lectin-like chaperone. Given the over-abundance of expression of this protein in epithelial layers selected from PP tissue and the role of this protein as a lectin-like chaperone, we propose that this protein is a valuable protein target to aid or facilitate the intracellular trafficking of antigens or antigens in particles following targeted delivery to M cells or PP tissue. Proteins comprising chimerics of calreticulin plus a polypeptide with an antigen of choice would therefore prove valuable in that regard.

Members of the 14-3-3 protein family have been identified as regulatory elements in intracellular signaling pathways and cell cycle control. There had been reports that 14-3-3 protein can be used as a marker for Creutzfeldt-Jacob Disease (CJD) in cerebrospinal fluid (CSF). It is proposed that this protein or the gene coding for it is valuable in the control of the M cell phenotype, and as a result it would be advantageous to co-deliver that protein or gene with a protein, antigen, or DNA vaccine.

Nucleoside diphosphate kinases (NDP kinases) form a family of oligomeric enzymes present in all organisms. Eukaryotic NDP kinases are hexamers composed of identical subunits (approximately 17 kDa). A distinctive property of human NDPK-B is its ability to stimulate gene transcription. This property is independent of its catalytic activity and is possibly related to the role of this protein in cellular events including differentiation and tumor metastasis. Given our discovery of the increased expression of nucleoside diphosphate kinase B in M cell enriched PP FAE cells, we propose the importance of this protein in determining or controlling M cell phenotype, in M cell development, and optimal activation or priming of the mucosal immune system.

Example 5

Gene expression analysis of rat PP and NPP tissue samples

In addition to the proteomic studies highlighted above, PP and NPP tissue samples were sent for gene expression analysis to CLONTECH Laboratories Inc. (a division of Becton Dickinson (BD) Biosciences) who then extracted RNA from the tissues to probe on ATLAS™1.2 rat arrays. The data containing differential expression levels of 1,200 genes many of which are presented in Table 1 below. The data show over-expression of many proteins. In Table 1, over-expressed genes are shown in bold and italicized.

In Table 1, "N/C" means not calculated due to manually-determined inconsistencies in one or both spots, and "?" means low confidence level (small difference).

Also, over-expressed genes from Table 1 that had a fold change above 0, as well as over-expressed genes are shown in Table 2 below with corresponding GenBank accession numbers for rat and human origin.

Based on the results (ratio PP/ Normal epithelial tissue) in Table 1, the following proteins are of the particular interest: clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, Ca²⁺-dependant phospholipase A2 precursor, ribosomal proteins S12, S11, L12, L11, S29, S19, L21, L19, L13, L44, and L36A.

In addition a series of genes coding for different TFs was noted including the following:

Jun-B; c-jun related TF,
 Jun-D; c-jun related TF,
 STAT 3 - signal transducer and activator of transcription 3,
 NF-kappa β Tf p105 subunit,

CREB active TF,
New england deaconess TF,
C-jun proto-oncogene; TF AP-1; RJG-9,
S-myc proto-oncogene; myc related,
5 Nm23-M2; nucleoside diphosphate kinase B; metastasis reducing protein,
NDK-B; nucleoside diphosphate kinase B ; metastasis reducing protein,
Lim-2; embryonic motor neuron topographic organizer; homeobox protein LIM-2, and
C-est-I proto-oncogene; p54.

10 TF coding genes such as these are considered here to be important in the
development of M cell phenotype and in priming the immune system. Their co-delivery or co-
targeting with DNA vaccine genes and/or with vaccines is expected to enhance activation of
mucosal immunity to the co-delivered DNA vaccine and/or antigen by virtue of their priming
of the cells to give a better mucosal immunity outcome.

15

TABLE 1 GENE EXPRESSION DATA FROM ATLAS 1.2 RAT ARRAY ANALYSIS									
#	coordinate	Spot Intensity		RATIO		Difference		GENE	
		PP1	NE1						
1	A03c	38	6	0.16	-32			T-cell surface glycoprotein CD5 precursor; lymphocyte glycoprotein LY1	
2	A03q	31	57	1.84	26			CD4 homologue, W3/25 antigen	
3	A04f	20	72	3.60	52			signal transducer CD24 precursor; heat stable antigen (HSA); necladrin	
4	A04i	17	40	2.35	23			CD2, membrane glycoprotein, T-cell marker	
5	A04j	8	34	4.25	26			scavenger receptor class B type I	
6	A04m	16	31	1.94	15			SR13 myelin protein; peripheral myelin protein 22 (PMP-22); CD25 protein	
7	A05a	4	17	4.25	13			glutamyl aminopeptidase A	
8	A05i	52	29	0.56	-23			I-kB (I-kappa B) alpha chain; RLIF-1 gene product	
9	A05m	6	18	3.00	12			interferon regulatory factor 1 (IRF1)	
10	A06a	7	21	3.00	14			LIM domain protein CLP36, homologous to rat RIL	
11	A06c	152	78	0.51	-74			Gax, growth-arrest-specific protein	
12	A07c	42	24	0.57	-18			G1/S-specific cyclin D3 (CCND3)	
13	A07i	17	3	0.18	-14			M-phase inducer phosphatase 2 (MIP2); cell division control protein 25 B (CDC25B)	
14	A07n	37	20	0.54	-17			p55cdc; cell division control protein 20	
15	A08e	56	24	0.43	-32			prothymosin-alpha (PTMA)	
16	A08n	32	105	3.28	73			antigen peptide transporter 1	
17	A08q	13	36	2.77	23			proteasome delta subunit precursor; macropain delta; multicatalytic endopeptidase complex delta;	
18	A09i	15	27	1.80	12			proteasome component C13 precursor; macropain subunit C13; multicatalytic endopeptidase	
19	A09j	341	893	2.62	552			apolipoprotein A-I precursor (APO-AI)	
20	A09k	72	723	10.04	651			apolipoprotein A-IV precursor (APO-AIV)	
21	A11i	1	13	13.00	12			ErbB3 EGF receptor-related proto-oncogene; HER3	
22	A12c	21	48	2.29	27			A-raf proto-oncogene	
23	A12i	19	45	2.37	26			rac-alpha serine/threonine kinase (RAC-PK-alpha); protein kinase B (PKB); AKT1	
24	A13k	208	60	0.29	-148			HSP64; Hsp60-beta; heat shock 60kD protein	
25	A14c	7	38	5.43	31			glutathione S-transferase Ya subunit (GST YA); ligandin subunit 1 alpha	
26	A14d	65	162	2.49	97			microsomal glutathione S-transferase (GST12; MGST1)	
27	A14e	94	160	1.70	66			glutathione S-transferase Yb subunit; GST subunit 4 mu (GSTM2)	
28	A14g	129	265	2.05	136			glutathione S-transferase P subunit; GST subunit 7 pl (GST7-7)	
29	A14n	7	19	2.71	12			NADPH-cytochrome P450 reductase (CPR); POR	
30	B01h	121	207	1.71	86			copper-zinc-containing superoxide dismutase 1 (Cu-Zn SOD1)	
31	B02f	4	25	6.25	21			fructose (glucose) transporter	
32	B05k	13	36	2.77	23			sodium channel SCN2B, beta 2 subunit, brain	
33	B06d	28	12	0.43	-16			potassium channel, inward rectifier 11	
34	B07k	2	18	9.00	16			proton-coupled dipeptide cotransporter	
35	B08b	59	181	3.07	122			fibroblast ADP/ATP carrier protein; ADP/ATP translocase 2; adenine nucleotide translocator 2	
36	B09i	7	43	6.14	36			sodium-glucose cotransporter 1	
37	B09l	40	115	2.88	75			Na+/K+ ATPase alpha 1 subunit	
38	B09n	43	78	1.81	35			vacuolar ATP synthase 16-kDa proteolipid subunit; ATP6C; MVP; ATP1	
39	B10b	10	78	7.80	68			sodium/potassium-transporting ATPase beta 1 subunit (ATP1B1)	
40	B12e	10	68	6.80	58			urate transporter/channel	
41	B12f	143	297	2.08	154			ATP synthase lipid-binding protein P1 precursor; ATPase protein 9; ATP5G1	
42	B12g	21	112	5.33	91			ATP synthase, subunit c, P2 gene	
43	B12m	18	38	2.11	20			annexin IV(ANX4); lipocortin IV; 36-kDa zymogen granule membrane-associated protein (ZAP36)	
44	B13f	37	127	3.43	90			lipocortin 2	
45	B14d	59	326	5.53	267			fatty acid-binding protein (liver, L-FABP); Z-protein; squalene- & steroid-carrier protein (SCP); P14	
46	B14f	49	157	3.20	108			fatty acid-binding protein (intestinal, I-FABP; FABPI)	

#	coordinate	Spot intensity	PP1	NET	RATIO	Difference	GENE
47	C02f	19	78	4.11	59	fructose-bisphosphate aldolase B (ALDOB); liver-type aldolase	
48	C02g	37	76	2.05	39	fructose-bisphosphate aldolase A (ALDOA); muscle-type aldolase	
49	C02i	15	30	2.00	15	testis fructose-6-phosphate 2-kinase/fructose 2,6-bisphosphate (testis 6PF-2-K/fru-2,6-P2ase); 6-	
50	C02n	604	1123	1.86	519	cytochrome c oxidase subunit Vb & Via precursor (COX5B)	
51	C03a	9	20	2.22	11	cytochrome B5 (CYB5)	
52	C03e	65	138	2.12	73	mitochondrial hydroxymethylglutaryl-CoA synthase precursor (HMG-CoA synthase); 3-hydroxy-3-	
53	C03f	177	760	4.29	583	cytochrome oxidase, subunit I, Sertoli cells	
54	C03g	27	47	1.74	20	ATPase, subunit F, vacuolar (vaf)	
55	C03i	641	1087	1.70	446	cytochrome c oxidase, subunit IV, mitochondrial	
56	C03j	46	147	3.20	101	cytochrome c oxidase, subunit Va, mitochondrial	
57	C03l	317	180	0.57	-137	cytochrome c oxidase, subunit VIII	
58	C04b	189	351	1.86	162	mitochondrial ATP synthase beta subunit precursor (ATP5B)	
59	C04g	23	127	5.52	104	creatine kinase, ubiquitous, mitochondrial	
60	C06c	6	52	8.67	46	fatty acid amide hydrolase	
61	C06j	73	175	2.40	102	cytochrome P450 17 (CYP17); P450C17; CYPXVII; sterol 17-alpha-hydroxylase/17,20 lyase	
62	C07i	9	51	5.67	42	cytochrome P-450 4F1, hepatic tumour	
63	C07l	19	39	2.05	20	cytochrome P-450 4F4	
64	C08a	17	33	1.94	16	cytochrome P-450 4F5	
65	C08i	8	24	3.00	16	adenylate kinase 3	
66	C08m	32	65	2.03	33	cAMP-dependent protein kinase type I-alpha regulatory chain	
67	C09e	7	30	4.28	23	glutathione synthetase (GSH synthetase; GSH-S; GSS); glutathione synthase	
68	C10e	25	10	0.40	-15	carbonic anhydrase 4	
69	C10i	19	9	0.47	-10	alkaline phosphatase	
70	C10k	21	5	0.24	-16	dopamine beta-hydroxylase	
71	C10l	29	11	0.38	-18	acetylcholinesterase, T subunit, glycolipid-anchored	
72	C10m	228	445	1.95	217	NADPH: alcohol dehydrogenase; aldehyde reductase (ALR); 3-dG-reducing enzyme	
73	C11b	19	33	1.74	14	calcium binding protein 2 (CABP2); endoplasmic reticulum stress protein (ERP72); protein disulfide	
74	C11d	97	55	0.57	-42	60S ribosomal protein L44; L36A	
75	C11e	1728	527	0.30	-1201	40S ribosomal protein S12	
76	C11g	400	179	0.45	-221	ribosomal protein L11	
77	C11h	670	355	0.53	-315	ribosomal protein L13	
78	C11i	987	424	0.44	-343	ribosomal protein L12	
79	C11k	812	413	0.51	-399	S19; 40S ribosomal protein S19	
80	C11l	554	286	0.52	-268	60S ribosomal protein L21	
81	C11m	1405	737	0.52	-668	60S ribosomal protein L19 (RPL19)	
82	C11n	315	109	0.35	-206	40S ribosomal protein S11	
83	C12b	158	48	0.30	-110	Fts-1; putative v-fos transformation effector protein; yeast mitochondrial protein import homolog; 40S	
84	C12d	1451	787	0.54	-664	elongation factor 2 (EF2)	
85	C12i	346	10	0.03	-336	clusterin (CLU); testosterone-repressed prostate message 2 (TRPM2); apolipoprotein J; sulfated	
86	C12m	14	0	Undefined	-14	activator of apoptosis harakiri (HRK); neuronal death protein 6 (DP6); BID3	
87	C13c	19	5	0.26	-14	SURVIVAL OF MOTOR NEURON (RSMN)	
88	D08l	6	17	2.83	11	retinoid X receptor alpha (RXR alpha; RXRA); NR2B1	
89	D12c	6	18	3.00	12	INOSITOL TRIPHOSPHATE RECEPTOR SUBTYPE 3	
90	E01m	10	50	5.00	40	neurotrophin 3 precursor (NTF3); neurotrophic factor; HDNF; nerve growth factor 2 (NGF2)	
91	E02c	19	8	0.42	-11	transforming growth factor, beta 1	
92	E03k	40	185	4.63	145	C-type natriuretic peptide precursor (CNP; NPPC)	
93	E04b	17	7	0.41	-10	thyroid stimulating hormone, beta	
94	E07c	33	15	0.45	-18	c-src-kinase (CSK) & negative regulator; tyrosine-protein kinase	

#	coordinate	Spot Intensity PP1	NE1	RATIO	Difference	GENE
95	E08a	105	249	2.37	144	extracellular signal-regulated kinase 1 (ERK1); mitogen-activated protein kinase 1 (MAP kinase 1; protein kinase C delta type (PKC-delta)
96	E08i	8	22	2.75	14	Insulin receptor-related receptor-alpha (sIRR-1)
97	E08j	36	68	1.89	32	CamK II; calcium/calmodulin-dependent protein kinase brain type II beta
98	E10f	3	17	5.67	14	CamK I; calcium/calmodulin-dependent protein kinase type I + CaM-like protein kinase
99	E10g	3	13	4.33	10	Casein kinase I delta; CKId; 49-kDa isoform
100	E10i	18	42	2.33	24	cyclin-dependent kinase 4 (CDK4); cell division protein kinase 4; PSK-J3
101	E11e	15	29	1.93	14	protein phosphatase 2C isoform; Mg2+ dependent protein phosphatase beta isoform
102	E12a	23	45	1.96	22	<i>guanine nucleotide-binding protein G(i) alpha 2 subunit (GNAI2); adenylate cyclase-</i>
103	E13i	40	19	0.46	-21	<i>rab12, ras related GTPase</i>
104	E14i	288	661	2.47	393	RalGDSB; GTP/GDP dissociation stimulator for a ras-related GTPase
105	F01e	10	29	2.90	19	<i>calcium-dependent phospholipase A2 precursor (PLA2); phosphatidylcholine 2-</i>
106	F02a	189	97	0.51	-92	<i>phospholipase C beta 3 (PLC-beta 3)</i>
107	F02h	15	79	5.27	64	14-3-3 protein zeta/delta; PKC inhibitor protein-1; KCIP-1; mitochondrial import stimulation factor
108	F04e	33	57	1.73	24	14-3-3 protein epsilon; PKC inhibitor protein-1; KCIP-1; mitochondrial import stimulation factor L
109	F04k	15	29	1.93	14	presenilin 1 (PSNL1; PSEN1; PS1); S182 protein
110	F05e	8	21	2.63	13	PDGF-associated protein
111	F05j	6	17	2.83	11	ADP-ribosylation factor 5 (ARF5)
112	F06a	59	143	2.42	84	dipeptidase (DPEP1)
113	F06j	12	22	1.83	10	granzyme M precursor (GZMM); MET-ASE; natural killer cell granular protease; RNK-MET-1
114	F07m	7	29	4.14	22	angiotensin converting enzyme (ACE; somatic; dipeptidyl carboxypeptidase I; kininase II
115	F08e	11	100	9.09	89	aminopeptidase B
116	F08h	18	48	2.67	30	kidney aminopeptidase M (APM)
117	F08j	39	92	2.36	53	metalloendopeptidase neprin beta subunit
118	F08k	19	81	4.26	62	endothelin converting enzyme
119	F08l	5	15	3.00	10	gelatinase A
120	F09a	3	16	5.33	13	cathepsin L
121	F09q	13	27	2.08	14	proteasome component C3
122	F09l	14	28	2.00	14	leukocyte common antigen-related tyrosine phosphatase (LAR)
123	F12a	6	16	2.67	10	ornithine decarboxylase (ODC)
124	G31	12	39	3.25	27	cytoplasmic beta-actin (ACTB)
125	G43	287	1132	3.94	845	<i>40S ribosomal protein S29 (RPS29)</i>
126	G47	7327	3614	0.49	-3713	

Table 2
RAT GENES (PP VS. NPP)

GENE	Fold change	GenBank ID Human	GenBank ID Rat
activator of apoptosis harakiri (HRK); neuronal death protein 5 (DP5); BID3	**	U76376.1	D83697
RET ligand 1 (RET1)	**	-	U97142
P2X purinoceptor 1; ATP receptor P2X1; purinergic receptor; RP-2 protein	**	P51575	U14414
leukocyte common antigen precursor (LCA); CD45 antigen; T200; PTPRC	**	Y00638	M10072
amphiphysin II (AMPH2)	**	AF001383.1	Y13380
Jak3 tyrosine-protein kinase; Janus kinase 3	**	XM_038595.3	D28508
DCC; netrin receptor; immunoglobulin gene superfamily member; former tumor suppressor protein candidate	**	M32292.1	AH002168.1
c-fgr proto-oncogene	**	AAA52762.1	X57018.1
small inducible cytokine A3 precursor (SCYA3); macrophage inflammatory protein 1 alpha precursor (MIP1-alpha; MIP1A)	**	P10147	AF119381.1
protein kinase C beta-I type (PKC-beta I) + protein kinase C beta-II type (PKC-beta II)	**	X06318	P04410
E-selectin precursor; endothelial leukocyte adhesion molecule 1 (ELAM-1); leukocyte-endothelial cell adhesion molecule 2 (LECAM2); CD62E	**	P16581	L25527
T-cell receptor CD3 zeta subunit	**	J04132.1	L08447.1
Protein kinase C-binding protein beta15; RING-domain containing	**	-	U48248
G1/S-specific cyclin C (CCNC)	**	AAC50825.1	D14013
maspin; protease inhibitor 5 (PI5); tumor suppressor	**	U04313.1	U58857
peptide/histidine transporter	**	-	AB000280
acetyl-CoA carboxylase (ACC); biotin carboxylase	**	X68968.1	AH002123.1
fibroblast growth factor receptor subtype 4	**	L03840.1	M91599
LCR-1; putative chemokine and HIV coreceptor homolog; G protein-coupled receptor	**	-	U54791
tumor necrosis factor alpha precursor (TNF-alpha; TNFA); cachectin	**	AF043342.1	X66539
CC chemokine MIP3 alpha exodus	**	-	U90447.1
luteinizing hormone, alpha	**	NM_000735.2	V01252
Ctk; non-receptor protein tyrosine kinase (batk)	**	P42679	L34542.1
RhoGAP; p122	**	-	S54293
Adenylyl cyclase type V	**	M83533.1	M96159
cathepsin S precursor (CTSS)	**	P25774	L03201.1
O-6-methylguanine-DNA methyltransferase (MGMT); methylated-DNA-protein-cysteine methyltransferase	**	M31767.1	NM_012861.1
clusterin (CLU); testosterone-repressed prostate message 2 (TRPM2); apolipoprotein J; sulfated glycoprotein 2 (SGP2); dimeric acid glycoprotein (DAG)	34.60	X14723	U02391.1
T-cell surface glycoprotein CD5 precursor; lymphocyte glycoprotein LY-1 (LYT1)	6.33	X04391.1	D10728
M-phase inducer phosphatase 2 (MPI2); cell division control protein 25 B (CDC25B)	5.67	S78187.1	D16237
dopamine beta-hydroxylase	4.20	Y00096.1	L12407

GENE	Fold change	GenBank ID Human	GenBank ID Rat
SURVIVAL OF MOTOR NEURON(RSMN)	3.80	AAC50473.1	U75369
HSP84; HSP90-beta; heat shock 90kD protein	3.47	XM_055551.3	S45392
Fte-1; putative v-fos transformation effector protein; yeast mitochondrial protein import homolog; 40S ribosomal protein S3A ; RPS3A	3.29	M84711.1	M84716.1
40S ribosomal protein S12	3.28	X53505	M18547
40S ribosomal protein S11	2.89	X06617	K03250
acetylcholinesterase, T subunit, glycolipid-anchored	2.64	M55040.1	X71089.1
carbonic anhydrase 4	2.50	NM_000717.2	I52551
thyroid stimulating hormone, beta	2.43	S70587.1	M13897.1
transforming growth factor, beta 1	2.38	M34057	NM_021578.1
prothymosin-alpha (PTMA)	2.33	AF257099.1	M20035
potassium channel, inward rectifier 11	2.33	-	D42145
ribosomal protein L12	2.28	L06505.1	X53504.1
ribosomal protein L11	2.23	X79234.1	X62146.1
c-src-kinase (CSK) & negative regulator; tyrosine-protein kinase	2.20	X59932.1	X58631
alkaline phosphatase	2.11	AAA98616.1	S18408
guanine nucleotide-binding protein G(i) alpha 2 subunit (GNAI2); adenylate cyclase-inhibiting G alpha protein	2.11	XM_041507.1	NM_031035.1
40S ribosomal protein S29 (RPS29)	2.03	NM_001032.2	X59051
S19; 40S ribosomal protein S19	1.97	P39019	P17074
Gax, growth-arrest-specific protein	1.95	-	Z17223.1
calcium-dependent phospholipase A2 precursor (PLA2); phosphatidylcholine 2-acylhydrolase (PLA2-10; PLA2G5)	1.95	M22430.1	U38376.1
60S ribosomal protein L21	1.94	P46778	M27905
60S ribosomal protein L19 (RPL19)	1.91	X63527	J02650
ribosomal protein L13	1.89	P26373	X78327.1
p53cdc; cell division control protein 20	1.85	AF099644.1	AF052695.1
elongation factor 2 (EF2)	1.84	X51466	Y07504.1
I-kB (I-kappa B) alpha chain; RL/IF-1 gene product	1.79	X63594.1	AF388201.1
60S ribosomal protein L44; L36A	1.76	M15661	P10661
cytochrome c oxidase, subunit VIIIh	1.76	J04823.1	NM_012786.1
G1/S-specific cyclin D3 (CCND3)	1.75	NM_001760.2	NM_012766.1
cytochrome c oxidase, subunit IV, mitochondrial	0.59	AF017115.1	X14209
glutathione S-transferase Yb subunit; GST subunit 4 mu (GSTM2)	0.59	-	X04229.1
copper-zinc-containing superoxide dismutase 1 (Cu-Zn SOD1)	0.58	-	NM_017050.1
14-3-3 protein zeta/delta; PKC inhibitor protein-1; KCIP-1; mitochondrial import stimulation factor S1 subunit	0.58	U28964.1	L07913.1
calcium binding protein 2 (CABP2); endoplasmic reticulum stress protein (ERP72); protein disulfide isomerase-related protein precursor	0.58	XM_012077.4	M86870
ATPase, subunit F, vacuolar (vaf)	0.57	AF047436.1	U43175.1
proteasome component C13 precursor; macropain subunit C13; multicatalytic endopeptidase complex subunit C13; PSMB8	0.56	P28062	NM_080767.1
vacuolar ATP synthase 16-kDa proteolipid subunit; ATP6C; MVP; ATP6L	0.55	NM_001695.1	M62762.1

GENE	Fold change	GenBank ID Human	GenBank ID Rat
dipeptidase (DPEP1)	0.55	NM_004413.1	M94056
CD4 homologue, W3/25 antigen	0.54	BC025782.	M15768.1
mitochondrial ATP synthase beta subunit precursor (ATP5B)	0.54	NM_001686.1	M19044.1
cytochrome c oxidase subunit Vb & VIa precursor (COX5B)	0.54	M59250.1	X14208.1
insulin receptor-related receptor-alpha (sIRR-1)	0.53	-	M90660.1
cyclin-dependent kinase 4 (CDK4); cell division protein kinase 4; PSK-J3	0.52	P11802	P35426
14-3-3 protein epsilon; PKC inhibitor protein-1; KCIP-1; mitochondrial import stimulation factor L subunit	0.52	XM_088041.1	D30739.1
SR13 myelin protein; peripheral myelin protein 22 (PMP-22); CD25 protein	0.52	-	M69139.1
cytochrome P-450 4F5	0.52	-	AF288818.1
NADP+ alcohol dehydrogenase; aldehyde reductase (ALR); 3-dG-reducing enzyme	0.51	J04794.1	D10854.1
protein phosphatase 2C isoform; Mg2+ dependent protein phosphatase beta isoform	0.51	-	S90449.1
testis fructose-6-phosphate 2-kinase/fructose 2,6-biphosphate (testis 6PF-2-K/fru-2,6-P2ase); 6-phosphofructo- 2-kinase; fructose-2,6-bisphosphatase	0.50	NM_002625.1	X15579.1
proteasome component C3	0.50	D00760	J02897.1
cAMP-dependent protein kinase type I-alpha regulatory chain	0.49	P10644	P09456
cytochrome P-450 4F4	0.49	-	U39206.1
fructose-bisphosphate aldolase A (ALDOA); muscle-type aldolase	0.49	XM_043948.2	NM_012495.1
glutathione S-transferase P subunit; GST subunit 7 pi (GST7-7)	0.49	-	X02904.1
ATP synthase lipid-binding protein P1 precursor; ATPase protein 9; ATP5G1	0.48	NM_005175.1	NM_017311.1
cathepsin L	0.48	M20496.1	Y00697.1
annexin IV(ANX4); lipocortin IV;36-kDa zymogen granule membrane-associated protein (ZAP36)	0.47	XM_031596.3	NM_024155.1
mitochondrial hydroxymethylglutaryl-CoA synthase precursor (HMG-CoA synthase); 3-hydroxy-3-methylglutaryl-CoA synthase; HMGCS2	0.47	P54868	P22791
cytochrome B5 (CYB5)	0.45	M22865.1	D13205.1
A-raf proto-oncogene	0.44	P10398	X06942
Casein kinase I delta; CKId; 49-kDa isoform	0.43	P48730	Q06486
CD2, membrane glycoprotein, T-cell marker	0.43	M14362.1	X05111.1
kidney aminopeptidase M (APM)	0.42	XM_087746.1	M26710
rac-alpha serine/threonine kinase (RAC-PK-alpha); protein kinase B (PKB); AKT1	0.42	P31749	Y15748.1
extracellular signal-regulated kinase 1 (ERK1); mitogen-activated protein kinase 1 (MAP kinase 1; MAPK1); insulin- stimulated microtubule-associated protein-2 kinase; MNK1; PRKM3; ERT2; p44-MAPK	0.42	P27361	P21708
cytochrome P450 17 (CYP17); P450C17; CYPXVII; steroid 17-alpha-hydroxylase/17,20 lyase	0.42	NM_000102.2	X69816.1
ADP-ribosylation factor 5 (ARF5)	0.41	NM_001662.	NM_024149.1

GENE	Fold change	GenBank ID Human	GenBank ID Rat
rab12, ras related GTPase	0.41	-	M83676.
microsomal glutathione S-transferase (GST12; MGST1)	0.40	XM_048886.3	J03752
apolipoprotein A-I precursor (APO-AI)	0.38	X02162	M00001
presenilin 1 (PSNL1; PSEN1; PS1); S182 protein	0.38	XM_007441.1	D82363
amonipectidase B	0.38	XM_087242.1	U61696
leukocyte common antigen-related tyrosine phosphatase (LAR)	0.38	-	U00477.1
NADPH-cytochrome P450 reductase (CPR); POR	0.37	S90469	NM_031576.1
protein kinase C delta type (PKC-delta)	0.36	NM_006254.1	M18330
proteasome delta subunit precursor; macropain delta; multicatalytic endopeptidase complex delta; proteasome subunit Y; proteasome subunit 5; PSMB6	0.36	X61971.1	NM_057099.1
sodium channel SCN2B, beta 2 subunit, brain	0.36	AAC05208.1	NM_012877.
retinoid X receptor alpha (RXR alpha; RXRA); NR2B1	0.35	XM_088424.1	NM_012805.1
PDGF-associated protein	0.35	U41745.1	U41744.1
Na+/K+ ATPase alpha 1 subunit	0.35	AAA51801.1	M28647
RaIGDSB; GTP/GDP dissociation stimulator for a ras-related GTPase	0.34	-	NM_019250.1
interferon regulatory factor 1 (IRF1)	0.33	XM_034862.1	M34253
LIM domain protein CLP36, homologous to rat RIL	0.33	AJ310549.1	U23769.1
adenylate kinase 3	0.33	XM_016642.3	NM_013218.1
INOSITOL TRIPHOSPHATE RECEPTOR SUBTYPE 3	0.33	-	L06096.1
endothelin converting enzyme	0.33	Z35307.1	D29683
fibroblast ADP/ATP carrier protein; ADP/ATP translocase 2; adenine nucleotide translocator 2 (ANT2)	0.33	J02683	D12771
cytochrome c oxidase, subunit Va, mitochondrial	0.31	M22760.1	X15030
fatty acid-binding protein (intestinal; I-FABP; FABPI)	0.31	M18079	M18080.1
ornithine decarboxylase (ODC)	0.31	X16277	D11372.1
antigen peptide transporter 1	0.30	X57522	P36370
lipocortin 2	0.29	D00017.1	S73557
signal transducer CD24 precursor; heat stable antigen (HSA); nectadrin	0.28	P25063	U49062
cytoplasmic beta-actin (ACTB)	0.25	M10277.1	V01217
fructose-bisphosphate aldolase B (ALDOB); liver-type aldolase	0.24	XM_042788.1	M10149
granzyme M precursor (GZMM); MET-ASE; natural killer cell granular protease; RNK-MET-1	0.24	NM_005317.2	Q03238
scavenger receptor class B type I	0.24	-	AB002151.1
glutamyl aminopeptidase A	0.24	XM_003595.2	S73583
metalloendopeptidase meprin beta subunit	0.23	-	M88601.1
glutathione synthetase (GSH synthetase; GSH-S; GSS); glutathione synthase	0.23	U34683.1	L38615.1
cytochrome oxidase, subunit I, Sertoli cells	0.23	S79304	S79304
CamK I; calcium/calmodulin-dependent protein kinase type I + CaM-like protein kinase	0.23	Q14012	L24907
C-type natriuretic peptide precursor (CNP; NPPC)	0.22	NM_024409.1	D90219

GENE	Fold change	GenBank ID Human	GenBank ID Rat
neurotrophin 3 precursor (NTF3); neurotrophic factor; HDNF; nerve growth factor 2 (NGF2)	0.20	M37763.1	M34643
phospholipase C beta 3 (PLC-beta 3)	0.19	NM_000932.1	M99567
ATP synthase, subunit c, P2 gene	0.19	D13119.1	D13124
gelatinase A	0.19	NM_004530.1	U65656
glutathione S-transferase Ya subunit (GST YA); ligandin subunit 1 alpha	0.18	NM_000852.2	K01932
creatine kinase, ubiquitous, mitochondrial	0.18	XM_016524.4	X59737
fatty acid-binding protein (liver; L-FABP); Z-protein; squalene- & sterol-carrier protein (SCP); P14	0.18	NM_001443.1	M35991
cytochrome P-450 4F1, hepatic tumour	0.18	-	NM_019623.1
CamK II; calcium/calmodulin-dependent protein kinase brain type II beta	0.18	NM_001220.1	M16112
sodium-glucose cotransporter 1	0.16	P13866	U03120
fructose (glucose) transporter	0.16	AAB60641	D13871.1
urate transporter/channel	0.15	-	U67958
sodium/potassium-transporting ATPase beta 1 subunit (ATP1B1)	0.13	NM_001677.1	NM_013113.1
fatty acid amide hydrolase	0.12	U82535.1	U72497
proton-coupled dipeptide cotransporter	0.11	-	D50306.1
angiotensin converting enzyme (ACE; somatic; dipeptidyl carboxypeptidase I; kinase II	0.11	NM_000789.1	NM_012544.1
apolipoprotein A-IV precursor (APO-AIV)	0.10	XM_052144.2	P02651
ErbB3 EGF receptor-related proto-oncogene; HER3	0.08	M29366.1	NM_017218.2
Jun-B; c-jun related TF,	**	M29039.1	X54686
S-myc proto-oncogene; myc related,	**		M29069
C-est-I proto-oncogene; p54.	5	AF193068.1	X55787.1
Jun-D; c-jun related TF,	1.79	X56681.1	D26307(mouse)
NF-kappaB Tf p105 subunit,	1.67	P19838	L26267.1
Nm23-M2; nucleoside diphosphate kinase B; metastasis reducing protein,	1.47		X68193.1 (mouse)
STAT 3 - signal transducer and activator of transcription 3,	1.16	NM_003150.1	NM_012747.1
CREB active TF,	1		M34356.1
New england deaconess TF,	1		U09229
Lim-2; embryonic motor neuron topographic organizer; homeobox protein LIM-2, and	1		L35572
NDK-B; nucleoside diphosphate kinase B ; metastasis reducing protein,	0.81		U29200.1
C-jun proto-oncogene; TF AP-1; RJG-9,	0.5	J04111.1	X17215.1

Symbols indicating fold changes in Table 2:

5

- ** : expressed in PP but not NPP, or in co-culture but not in Caco-2 cells.
- * : expressed in co-culture but not in Caco-2 cells (only repeated once).
- : expressed in Caco-2 but not in co-culture.

Example 5ATLAS array data on co-culture of human Caco-2 cells and Raji B-cells

5 In order to facilitate the routine study of M cell biology, there was a desire to establish
a suitable and representative in-vitro model. In the work carried out by Kernéis *et al.*
(Kerneis S, Caliot E, Stubbe H, Bogdanova A, Kraehenbuhl J, Pringault E (2000). Molecular
studies of the intestinal mucosal barrier physiopathology using co-cultures of epithelial and
immune cells: a technical update. *Microbes Infect* 2000 Jul;2 (9):1119-24), it was reported
10 that Peyer's patch lymphocytes co-cultured with Caco-2 cells trigger the phenotypic
conversion of enterocytes into cells that express morphological and functional M-cell
properties. This work was further developed by Gullberg *et al.* (Gullberg E, Leonard M,
Karlsson J, Hopkins AM, Brayden D, Baird AW, Artursson P. Expression of specific markers
and particle transport in a new human intestinal M-cell model. *Biochem Biophys Res*
15 *Commun* 2000 Dec 29; 279(3): 808-13) to create a simplified in vitro model of the human M-
cell. Co-cultures of physically separated human intestinal epithelial Caco-2 cells and B-cell
lymphoma Raji cells were established. The co-cultures were characterized under the criteria
of morphology, integrity, expression of M-cell markers and cell adhesion molecules (CAMs),
and altered particle transport. Using this construct, the epithelial cells were transformed to
20 cells with an M-cell-like morphology and had altered expression of potential human M-cell
markers (alkaline phosphatase down-regulation and Sialyl Lewis A antigen up-regulation).
The expression of intercellular adhesion molecule-1 and vascular cell adhesion molecule
was altered, and there was an increased binding of lectins wheat germ agglutinin and peanut
agglutinin with a 40-fold increase in microparticle transport. The particle transport was size-
25 dependent and could be inhibited at 4°C or by replacing the Raji B-cells with Jurkat T-cells.
Thus the comparison of RNA isolated from co-cultured Caco2 cells to that isolated from
normal Caco2 cells was designed to simulate a comparison of M cell RNA to normal gut
enterocyte RNA.

30 Isolation of total RNA from Co-Cultured Caco-2 cells

Caco-2 cell culture

Caco-2 cells were cultured in Dulbecco's Modified Eagles Medium (DMEM), 4.5g/L
glucose supplemented with 1% Mem, 10% FCS and 1% penicillin/streptomycin at 37°C and

5 5% CO₂ in 95% relative humidity. Cells were grown and expended in Falcon culture flasks and passaged once they attained 100% confluence. Caco-2 cells were seeded on Transwell Clear filters (Costar, 12mm diameter, 3.0um pore size)) at a density of 5x10⁵ cells/cm² and incubated in a 12 well culture plate with a medium change every second day. 1.0ml was added to the basolateral side and apical sides.

Raji cell culture

10 Raji B-lymphoma cells were cultured in RPMI 1640 Medium, with 1% (v/v) non-essential amino-acids, 10% FCS and 1% penicillin/streptomycin, 1% L-glutamine at 37°C and 5% CO₂ in 95% relative humidity. Cells were grown in suspension in Falcon tissue culture flasks and passaged by dilution every 5-7 days.

Co-culture: day 14 (treating with Raji B-cells)

15 After 14 days of culturing Caco-2 monolayers, 15-20ml of Raji cells were removed from the T75 flask and placed in a 20ml universal. The cells were centrifuged at 1000 rpm for 3 min. Cells were re-suspended at a concentration of 1x10⁶ cells/ml. 1ml of fresh complete DMEM was added to the apical and basolateral sides of the Caco-2 monolayer filters. 0.5 ml of 1x10⁶ Raji cells/ml cells was added to the basolateral side of the filters. For control filters (non co-culture) 0.5ml of Raji medium only was added to the basolateral side.

20

Isolation of Total RNA from co-cultured Caco-2 cells

25 After 4 days of co-culture the filters were rinsed in PBS. 0.5 ml of PBS was added to the apical side of each filter and the Caco-2 cells were scraped off the filter surface into suspension in the PBS. The cells from all the co-cultured Caco-2 filters were pooled, centrifuged at 1000 rpm for 3 min, the supernatant PBS was removed and the pellet was used for RNA extraction.

Analysis of mRNA expression

30 Total cellular RNA was extracted using an acid guanidinium thiocyanate-phenol-chloroform method. RNA's integrity was confirmed by gel electrophoresis and ethidium bromide staining. mRNA was reverse transcribed in the presence of P³² dATP, and the transcribed cDNA was purified by chromatography before being hybridized over night to the array membrane. Membranes were exposed to x-ray film using an intensifying screen for 3 days and the mRNA expression levels were analyzed by scanning the films with a

densitometer. Expression levels were normalized relative to internal standards, and relative increases in mRNA levels in co-cultured cells versus monoculture controls were calculated. Two hybridization experiments were performed using mRNA from two separate cell harvests. Results from the two experiments were pooled, and a summary of the findings was tabulated in Tables 3(a)-3(f). The identified genes are from the following groups: oncogenes, tumor suppressor genes, genes involved in the cell cycle, ion channels and transport, stress response genes, modulators and effectors, genes involved in intracellular transduction, genes linked to apoptosis, DNA synthesis, repair & recombination, transcription factors, DNA binding proteins, receptors, cell surface antigens, genes involved in cell adhesion, growth factors, cytokines, chemokines and hormones.

In Table 3, genes which were found to be exclusively over-expressed in the co-culture and not in the control Caco-2 monolayer are represented by **. A single asterisk represents genes that also were expressed in the co-culture and not in the control Caco-2 monolayer. However, these particular genes have been distinguished from the genes labeled with two asterisks as they were not expressed in both hybridization experiments performed, and will require confirmation in the future by PCR so as to rule out false positives/negatives. Genes not expressed in the co-culture but expressed in the Caco-2 monolayer controls are indicated by a minus symbol, "-".

Table 3 a: Oncogenes, Tumor Suppressors, Cell Cycle Regulators

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
Myeloid cell nuclear differentiation antigen (MNDA)	*	M81750
G1/S-specific cyclin D1 (CCND1); cyclin parathyroid adenomatosis 1 (PRAD1); bcl-1 oncogene	*	X59798
cyclin-dependent kinase 4 inhibitor 2 (CDK4I; CDKN2); p16-INK4; multiple tumor suppressor 1 (MTS1)	*	L27211.1
cyclin-dependent kinase inhibitor 1C (CDKN1C); p57-KIP2	*	U22398
ezrin; cytovillin 2; villin 2 (VIL2)	1.69	X51521
proto-oncogene tyrosine-protein kinase kit; c-kit; mast/stem cell growth factor receptor precursor(SCFR); CD117 antigen	1.55	L04143.1
proliferating cell nucleolar antigen P120; NOL1	1.52	M32110
jun proto-oncogene; avian sarcoma virus 17 oncogene homolog; transcription factor AP-1	1.47	J04111
C-src proto-oncogene (SRC1)	1.35	X59932
CDC-like kinase 3 (CLK3)	1.35	L29220
cell division cycle protein 25 nucleotide exchange factor (CDC25)	1.34	M91815.1
prothymosin alpha (PROT-alpha; PTMA)	1.32	M26708
40S ribosomal protein S19 (RPS19)	1.31	M81757
avian myelocytomatosis viral oncogene homolog (MYC)	1.30	V00568
CDC-like kinase 1 (CLK1)	1.27	L29219.1
cyclin-dependent kinase 4 inhibitor 2D (CDKN2D); p19-INK4D	0.69	U49399.1
vascular endothelial growth factor receptor 1 (VEGFR1); tyrosine-protein kinase receptor flt + soluble VEGFR; tyrosine-protein kinase receptor SFLT	0.62	XM_039993.2
neogenin	-	U61262.1
webB2 receptor protein-tyrosine kinase; neu proto-oncogene; c-erbB2 + HER2 receptor	-	M11730.1
N-ras; transforming p21 protein	-	AAA60255

Table 3 b: Ion Channels, Modulators, Effectors

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
extracellular signal-regulated kinase 3 (ERK3); mitogen-activated protein kinase 6 (MAP kinase 6; MAPK6; PRKM6); p97-MAPK	**	X14798.1
40-kDa heat-shock protein 1 (HSP40); DNAJ protein homolog 1 (HDJ1; DNAJ1)	**	D49547
70-kDa heat shock protein 1 (HSP70.1; HSPA1)	**	M11717
glutaredoxin	**	X76648
tyrosine kinase receptor tie-1 precursor	*	AAB84296
ras-related protein RAB3B	*	NM_002867.1
macMARCKS; MARCKS-related protein (MRP); MLP	*	P49006
mitogen-activated protein kinase 3 (MAPK3; PRKM3);	*	P27361

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
MAPK1; extracellular signal-regulated kinase 1 (ERK); microtubule-associated protein 2 kinase; insulin-stimulated MAP2kinase		
mitogen-activated protein kinase 9 (MAP kinase 9; MAPK9; PRKM9); c-jun N-terminal kinase 2 (JNK2); JNK55	*	NM_002752.1
60-kDa heat shock protein (HSP60); HSPD1; 60-kDa chaperonin; mitochondrial matrix protein P1 precursor; p60 lymphocyte protein; HUCHA60; GROEL	*	M22382.1
serine kinase	2.24	U09564.1
transferrin receptor (TFRC); CD71 antigen	1.80	M11507.1
Neurotrophic tyrosine kinase receptor-related 3: TKT precursor	1.63	U55017.1
phospholipase C (PLCL)	1.62	X14034.
cAMP-response element binding protein (CREB)	1.59	M27691.1
ephrin type-A receptor 1 precursor; tyrosine-protein kinase receptor eph	1.55	M18391
27-kDa heat-shock protein (HSP27); stress-responsive protein 27 (SRP27); estrogen-regulated 24-kDa protein; HSPB1	1.42	X54079.1
tyrosine kinase tnk1	1.42	XM_012654.3
ras-related protein RAB3A	1.38	XM_054457.2
janus kinase 3 (JAK3); leukocyte janus kinase (L-JAK)	1.33	XM_038595.3
dual-specificity mitogen-activated protein kinase kinase 1 (MAP kinase kinase 1; MAPKK 1; MKK1); extracellular signal-regulated kinase 1; ERK activator kinase 1	1.29	NM_002755.2
calcium/calmodulin-dependent protein kinase type IV catalytic subunit (CAMK IV); CAM kinase-GR	1.27	NM_001744.1
ras-related protein RAB5A	0.75	XM_053461.2
colon carcinoma kinase 4 precursor (CCK4) + transmembrane receptor PTK7	0.68	U33635.1
epithelial discoidin domain receptor 1 precursor (EDDR1; DDR1); cell adhesion kinase (CAK); TRKE; RTK6; protein tyrosine kinase 3A (PTK3A); neuroepithelial tyrosine kinase (NEP)	0.63	XM_004559.5
ras-related protein RAB6	0.27	M28212.1
cAMP-dependent protein kinase type I beta regulatory subunit (PRKAR1B)	0.23	M65066.1
tyrosine-protein kinase ack	-	CAC15525
T-lymphocyte maturation-associated protein MAL	-	P21145
orphan hormone nuclear receptor	-	U04897.1
LIM domain kinase 1 (LIMK-1)	-	P53667
protein kinase C alpha polypeptide (PKC-alpha; PKCA)	-	NM_002737.1
dual specificity mitogen-activated protein kinase kinase 3 (MAP kinase kinase 3; MAPKK3; MKK3); ERK activator kinase 3; MAPK/ERK kinase 3 (MEK3)	-	P46734
Yamaguchi sarcoma viral-related oncogene homolog; tyrosine -protein kinase lyn	-	M16038.1
protein-tyrosine phosphatase 1E	-	U12128.1

Table 3.c: Apoptosis, DNA Synthesis, Repair & Recombination

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
ubiquitin-conjugating enzyme E2 17-kDa (UBE2A); ubiquitin-protein ligase; ubiquitin carrier protein, HR6A	**	NM_003336.1
growth arrest & DNA-damage-inducible protein 153 (GADD153); DNA-damage-inducible transcript 3 (DDIT3); C/EBP homologous protein (CHOP)	**	S40706.1
growth factor receptor-bound protein 2 (GRB2); ASH protein	*	M96995.1
glutathione S-transferase A1 (GTH1; GSTA1); HA subunit 1; GST-epsilon	*	M21758.1
cytoplasmic dynein light chain 1 (HDLC1); protein inhibitor of neuronal nitric oxide synthase (PIN)	*	U32944.1
xeroderma pigmentosum group G complementing protein (XPG); X-ray repair-complementing defective repair in Chinese hamster cells 5 (XRCC5)	*	NM_021141.2
xeroderma pigmentosum group D complementing protein (XPD); X-ray repair-complementing defective repair in Chinese hamster cells 2 (XRCC2)	*	AF035587.1
RAD23 homolog A (RAD23A; hHR23A)	*	NM_005053.1
ataxia telangiectasia (ATM)	*	AAB38309
apoptosis regulator bcl-x	1.60	Z23115.1
caspase 9 precursor (CASP9); ICE-like apoptotic protease 6 (ICE-LAP6); apoptotic protease MCH6; apoptotic protease activating factor 3 (APAF3)	1.42	AB020979.1
CD40 receptor-associated factor 1 (CRAF1)	1.39	U21092.1
SL cytokine precursor; FMS-related tyrosine kinase 3 ligand (FLT3 ligand; FLT3LG)	1.35	NM_001459.1
cytochrome P450 reductase	1.33	AAD45961.1
X-ray repair complementing defective repair in Chinese hamster cells 1 (XRCC1)	1.25	M36089
Ku (p70/p80) subunit; ATP-dependent DNA helicase II 86-kDa subunit; lupus ku autoantigen protein; thyroid-lupus autoantigen (TLAA); CTC box binding factor 85-kDa subunit (CTCBF; CTC85); nuclear factor IV	0.74	X57500.1
caspase 10 precursor (CASP10); ICE-LIKE apoptotic protease 4 (ICE-LAP4); apoptotic protease MCH4; fas-associated death domain protein; interleukin 1 beta-converting enzyme 2 (FLICE2);	0.45	Q92851
inhibitor of apoptosis protein 2 (HIAP2; IAP2) + IAP homolog B; TNFR2-TRAF signaling complex protein 2; MIHB	-	Q13490
recA-like protein HsRad51; DNA repair protein RAD51 homolog	-	BAA02962.1
DNA damage repair & recombination protein 52 (RAD52)	-	B56529
DNA ligase III (LIG3); polydeoxyribonucleotide synthase	-	CAA59230.1

Table 3 d: Transcription Factors, DNA Binding Proteins

Gene	Fold change	GenBank ID
transcriptional activator hSNF2-alpha	**	D26155.1
early growth response protein 1 (EGR1); transcription factor ETR103; KROX24; zinc finger protein 225 (ZNF225); AT225	2.71	M62829.1
homeobox A1 protein (HOXA1); HOX1F	2.17	U10421.1
transcription factor NF-ATc	1.67	U08015.1
R kappa B DNA-binding protein	1.66	U08191.1
transcription initiation factor IID 31-kDa subunit (TFIID); TATA-box-binding protein-associated factor RNA polymerase II G 32-kDa subunit (TAFII32; TAF2G); TAFII31	1.57	M55654
homeobox protein hLim1; LHX1	1.51	NM_005568.1
helix-loop-helix protein HLH 1R21; DNA-binding protein inhibitor Id-3; HEIR-1	1.49	X69111.1
guanine nucleotide-binding protein G-s alpha subunit (GNAS); adenylate cyclase-stimulating G alpha protein	1.46	NP_000507.1
CCAAT-binding transcription factor subunit B (CBF-B); NF-Y protein subunit A (NF-YA); Hap2; CAAT-box DNA-binding protein subunit A	1.45	AAA40889.1
transcription factor LSF	1.37	B53771
homeobox 2.1 protein (HOX2A); HOXB5; HU1; HHO.C10	1.35	M92299.1
endothelial transcription factor GATA2	1.34	M68891.1
transcription factor Sp1 (TSFP1)	1.30	XM_028606.2
transcription factor ZFM1	0.62	G02919
zinc finger protein 161 (ZNF161); putative transcription activator DB1	0.26	NP_009077.1
stem cell protein (SCL); T-cell leukemia/lymphoma-5 protein (TCL5); T-cell acute lymphocytic leukemia-1 protein (TAL1)	-	AAA36598.1
neural retina-specific leucine zipper protein (NRL)	-	NP_006168
MSX-1 homeobox protein; HOX7	-	P28360
basic transcription factor 62-kDa subunit (BTF2)	-	AAA58399.1
paired box homeotic protein (PAX8) isoforms 8A/8B + isoforms 8C/8D	-	BAB59039.1
brain-specific homeobox/POU domain protein 3A (bm-3A); RDC-1; octamer binding transcription factor 1 (OTF1)	-	AAA65605.1
transcription factor E2-alpha (E2A); immunoglobulin enhancer binding factor E12; transcription factor-3 (TCF3)	-	AAA61146.1
transcriptional enhancer factor (TEF1); protein GT-IIC; transcription factor 13 (TCF13)	-	P28347
thioredoxin peroxidase 2 (TDPX2); thioredoxin-dependent peroxide reductase 2; proliferation-associated gene (PAG); natural killer cell enhancing factor A (NKEFA)	-	Q06830

Table 3 e: Receptors, Cell Surface Antigens, Cell Adhesion

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
interleukin-2 receptor gamma subunit (IL-2R gamma; IL2RG); cytokine receptor common gamma chain precursor; p64	*	AAA59145.1
interferon gamma receptor (IFNGR)	*	NM_000416.1
interleukin-1 receptor type I precursor (IL-1R1); IL-1R-alpha; p80; CDW121A antigen	*	M27492.1
neural-cadherin precursor (N-cadherin; NCAD); cadherin 2 (CDH2)	*	L34059
neural cell adhesion molecule L1 precursor (N-CAM L1); MIC5	*	M77640
integrin alpha 3 (ITGA3); galactoprotein B3 (GAPB3); very late antigen 3 alpha subunit (VLA3 alpha); CD49C antigen	*	M59911.1
leukocyte adhesion glycoprotein p150, 95 alpha subunit precursor; leukocyte adhesion receptor p150, 95; CD11C antigen; leu-M5; integrin alpha X (ITGAX)	*	M81695.1
integrin beta 4 (ITGB4); CD104 antigen	*	X51841.1
CD44 antigen precursor (CD44); phagocytic glycoprotein I (PGP1); HUTCH I; extracellular matrix receptor III (ECMR III); gp90 lymphocyte homing/adhesion receptor (LHR); hermes antigen; hyaluronate receptor; heparan sulfate proteoglycan; epican	1.51	XP_030326.1
glutamate receptor subunit epsilon 3 precursor (GRIN2C); N-methyl D-aspartate receptor subtype 2C (NMDAR2C; NR2C)	1.44	NP_000826.1
CD27L antigen receptor precursor; tumor necrosis factor receptor superfamily member 7 (TNFRSF7); T14	0.7	P26842
integrin alpha L (ITGAL); leukocyte adhesion glycoprotein alpha subunit precursor; leukocyte function-associated molecule 1 alpha chain (LFA1); CD11A antigen	0.45	P20701
interleukin 2 receptor alpha subunit precursor (IL-2 receptor alpha subunit; IL2RA); TAC antigen; CD25 antigen	0.41	P01589
CDW40 antigen; CD40L receptor precursor; nerve growth factor receptor-related B-lymphocyte activation molecule	0.35	CAA43045.1
granulocyte colony stimulating factor receptor precursor (GCSF-R); CD114 antigen	-	Q99062
low-affinity nerve growth factor receptor (NGF receptor; NGFR); GP80-LNGFR	-	AAB59544.1
neuromedin B receptor (NMBR); neuromedin-B-preferring bombesin receptor	-	NP_002502.1
granulocyte-macrophage colony-stimulating factor receptor alpha (GM-CSFR-alpha); CSW116 antigen	-	Q00941
platelet membrane glycoprotein IIIa precursor (GP3A); integrin beta 3 (ITGB3); CD61 antigen	-	P05106
integrin alpha 7B precursor (IGA7B)	-	CAA52348.1

Table 3 f: Growth Factors, Cytokines, Hormones

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
Interleukin-10 precursor (IL-10); cytokine synthesis inhibitory factor(CSIF)	**	M57627
Granulocyte-macrophage colony stimulatng factor (GM-CSF); CSF2	*	AAA52578.1
FMLP-related receptor I (FMLPRII); RMLP-related receptor I (RMLPRI)	*	AAA58482.1
Glia maturation factor beta (GMF-beta)	*	P17774
Hepatoma-derived growth factor (HDGF)	*	P51858
Macrophage inflammatory protein 1 alpha precursor (MIP1-alpha); tonsillar lymphocyte LD78 alpha protein; G0S19-1 protein; PAT 464.2; SIS-beta; small inducible cytokine A3 (SCYA3)	*	P10147
Monocyte chemotactic protein 1 precursor (MCP1); monocyte chemotactic and activating factor (MCAF); monocyte secretory protein JE; monocyte chemoattractant protein 1; HC11; small inducible cytokine A2 (SYCA2).	*	P13500
Oncostatin M (OSM)	*	NP_065391.1
Renin-binding protein (RENBP; RNBP)		XP_013053.3
Calgranulin B (CAGB); migration inhibitory factor-related protein 14 (MRP14); leukocyte L1 complex heavy chain; S100 calcium binding protein A9 (S100A9)	1.49	B31848
Placenta growth factors 1+2 (PLGF1 + PLGF2)	1.42	CAA38698.1
Vascular endothelial growth factor precursor (VEGF); vascular permeability factor (VPF)	1.42	AAA35789.1
Hepatocyte growth factor activator (HGF activator)	1.40	BAA74450.1
Follistatin-related protein precursor	1.34	AAA66062.1
Hepatocyte growth factor-like protein; macrophage stimulating protein (MSP)	1.29	AAA59872.1
Interferon gamma precursor (IFN-gamma, IFNG); immune interferon	1.29	P01579
WSL protein + TRAMP + Apo-3 + death domain receptor 3 (DDR3)	0.69	AAB41432.1
Neurotrophin-4 (NT4)	0.68	AAA60154.1
Interleukin-13 precursor (IL-13); NC30	0.39	P35225
Small inducible cytokine A5 (SYCA5); regulated on activation normal T-cell-expressed & secreted protein precursor (RANTES); SIS delta	0.38	XP_035842.1
Estrogen sulfotransferase (STE; EST1)	-	CAA72079.1
Keratinocyte growth factor (KGF); fibroblast growth factor 7 (FGF7)	-	AAA63210.1
Endothelial-monocyte activating polypeptide II (EMAP II)	-	AAA62202.1
Leukemia inhibitory factor precursor (LIF); differentiation-stimulating factor (D factor); melanoma derived LPL inhibitor (MLPLI); HILDA	-	B36282
Acidic fibroblast growth factor (AFGF) + heparin-binding growth factor 1 precursor (HBGF-1) + beta-endothelial cell growth factor (ECGF-beta)	-	AAA51672.1
Insulin-like growth factor-binding protein 3 precursor (IGF-binding protein 3; IGFBP3; IBP3)	-	P17936

Symbols (Fold Changes)

- 5 ** : Expressed in PP but not NPP, or in Co-culture but not Caco2.
 * : Expressed in Co-culture but no Caco2 (only repeated once)
 - : Expressed in Caco2 but not in co-culture.

Immunity

10 The events of the cell cycle occur under normal circumstances in a fixed sequence. Traditionally, the cycle is divided into two stages: cell division and the interphase. Cell division or mitosis is followed by cytokinesis and together they constitute the 'M phase' of the cell cycle. The interphase is divided up into the S, G₁ and G₂ phases. Briefly, during the S phase, DNA is replicated in preparation for mitosis, while the intermediate G phases are transitional periods involved in protein synthesis and cell growth. Activation of regulatory genes that control and maintain a cell's proliferative state by intracellular signals (discussed below) stimulates proliferation of the cell and initiates cell growth. A number of genes involved in these processes were differentially expressed in the co-culture model (as estimated by relative mRNA abundance) and discussed below.

15 The epithelial cells of the gut play an important part of the innate and specific immunity. IEC's are considered to be in a continuous controlled state of "physiological" inflammation and active processes continually take place to ensure that the tone of immunosuppression is maintained (Mayer, 2000). These unique regulators appear to control the mucosal immune system's condition. These distinct factors govern the immune response, whether it's immune suppression/tolerance, inflammation or a systemic immune response. A clearer understanding of the immunoregulatory features involved in mucosal immunity is clearly desirable and may lead to new approaches in disease and drug therapy. Genes detected in the co-culture model that may be related to or are involved in immune function in GALT are discussed below.

25 The gamma subunit of IL-2 receptor plays a pivotal role in formation of the full-fledged IL-2 receptor (Di Santo *et al.*, 1995). In an interesting study where infant rats were studied from pre- to post weaned life Masjedi *et al.* (1999) assessed alterations in expression and phenotype of cells in the gut-associated lymphoid tissue. At an age when the immune system is believed to be immature and functionally naive they discovered interleukin-2 receptor (IL-2R) expression peaked approximately four-fold at midweaning in Peyer's patches, compared with adult animals (day 70) suggesting that IL-2R expression is an adaptation to the host's environment. In a similar way, the presence of IL-2R specific for cells in the co-culture could be a direct result of the environment. The common gamma c

chain of the interleukin 2 receptor, gamma is also a component of the receptors for IL-4, IL-7, and IL-9 and plays a critical role in lymphoid development through its participation in the receptors for IL-2, IL-4, IL-7, IL-9, and IL-15 (Di Santo *et al.*, 1995)

Interferon- γ (IFN- γ) exhibits various properties including antiproliferative activity in neoplastic and normal cells, and regulatory roles in immune responses (Tsuiji *et al.*, 1998). Kjerrulf *et al.* (1997) found that in IFN- γ receptor knockout mice (IFN- γ R^{-/-}) reduced mucosal antibody responses and decreased Th1 and Th2 activity after oral immunization. The presence of IFN- γ receptor in the M cell co-culture model could possibly augment a cross-regulation between the two Th subsets in the gut mucosa. It is noteworthy that the ligand, IFN- γ , mRNA was increased in the co-culture that was supported further by the significant secretion of IFN- γ from co-culture monolayers.

The C-C chemokines macrophage inflammatory protein 1 (MIP1 α) and monocyte chemoattractant protein (MCP1) are synthesized and expressed by epithelial cells (Vainer *et al.*, 1998; Kolios *et al.*, 1999). The purpose of these chemokines expression in the co-culture model could be to function not only in leukocyte migration, but also as adhesins in the interaction between leukocytes and colonic epithelium. However, another C-C chemokine, RANTES, mRNA was observed to be reduced in the co-culture. The reasons for this are unclear. Perhaps, the chemoattractant activities of other chemokines such as IL-8, MIP1 α and MCP1 are sufficient for the M cell and in the absence of T cells the need for RANTES is not required.

From a gene delivery perspective, a higher capacity for translation and protein synthesis in PP tissue indicates that PP tissue is a preferred tissue to which to deliver genes coding for DNA vaccines or antigens. Thus the proposed higher translational capacity of PP tissue has implications for gene delivery especially DNA vaccine delivery and correspondingly antigen expression and local presentation to the mucosal immune system in the gastrointestinal tract. The TF coding genes may be important in priming M cells or precursor cells to M cells to adopt M cell phenotype and/or to facilitate priming of M cells to give a better immune cell outcome.

M cell receptors identified in Table 3(e) above are of particular interest in that they can be used for vaccine and delivery.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of an IL-2 receptor, a gamma c chain of an IL-2 receptor, interferon - γ , and a C-C chemokine.

Proliferation and Growth

Cyclin D1 is a protein involved in regulation of the cell cycle. Over-expression of the protein is associated with abnormal growth or neoplasia. This protein is positively induced by the p42/p44 MAP kinases (Lavoie *et al.*, 1996). It would be interesting if the neoplasia seen in M cells resulted from activation of this protein considering the coincidental induction of the p44 MAP kinase (ERK1) below. The reduction in cyclin-dependent kinase 4 inhibitor 2D (CDKN2D) mRNA that normally inhibits cell cycle progression (Guan *et al.*, 1996) would insinuate a similar function in the proliferation of these 'M cells.'

In contrast, the induction of cell cycle inhibitors such as cyclin-dependent kinase inhibitor (CDKI) and cyclin-dependent kinase 4 inhibitor (CDK4J) would appear to be working to counterbalance proliferative stimuli present in the M cell.

PLC-L (phospholipase C-deleted in lung carcinoma) is a putative tumor suppressor gene. It is believed that irregular (in fact deletion) expression of the PLC-L gene contributes to the growth of human lung carcinoma (Kohno *et al.*, 1995). It is possible then that its upregulation in the M cell model is acting as a negative regulator of growth in the cells, counterbalancing the many proliferative signals present.

Growth factor receptor-bound protein 2, GRB2, involved in growth factor control of ras signalling (Lowenstein *et al.*, 1992).

The intracellular signaling pathways responsible for cell cycle arrest and establishment of differentiated cells along the gut axis remain largely unknown particularly in the case for the development of M cells and the FAE. ERK3/MAPK6 is expressed solely in the co-culture. Extracellular signal-regulated kinases-1 (ERK1) also known as the p44 mitogen-activated protein (MAP) kinase (p44mapk) is also induced specifically in the co-culture model. ERK1 and ERK3 are proline-directed serine/threonine kinases that are activated in response to a variety of extracellular signals, including growth factors, hormones and neurotransmitters. These MAP kinases are key molecules involved in intracellular signal transduction, and are key regulators of cell proliferation in mammalian cells (Davis, 1995). Results indicate that elevated p42/p44 MAPK activities stimulate cell proliferation of intestinal cells, whereas low sustained levels of MAPK activities have correlated with cell cycle arrest and an increased expression of sucrase isomaltase (Aliaga *et al.*, 1999). It is tempting to speculate that the presence of ERK3 together with the other MAP kinases apart from their proliferative effects are in part responsible for a reduction in sucrase isomaltase, a characteristic effect in M cells.

Lying upstream in the ERK signal cascade the tyrosine/threonine protein kinase, MAPK kinase (MAPKK1) is implicated in the regulation of cell growth and differentiation through the activation of ERK. In addition it is interesting to note that MAPKK3 was deleted in the co-culture cells. MAPKK3 phosphorylates and activates p38 MAP kinase alpha and gamma isoforms (Enslen *et al.*, 1998). The induction of the MAPKK1 gene along with serine kinase coincides with the induction of ERK1, highlighting the ERK cascade as an important signalling cascade in M cell maintenance. It is interesting to note that ERK activation is responsible for terminal differentiation of components of the crypt-villus. (Taupin and Podolsky, 1999)

However, glia maturation factor- β (GMF- β) is potentially offsetting the ERK cascade effects. It is known to inhibit MAP kinases particularly ERK1 and ERK2 and yet promotes the p38 MAPK (Zaheer and Lim, 1996 and 1998).

Findings suggest that positive and negative regulation of MAPK activity are associated with loss of normal growth control and may be involved in carcinogenesis of colon cancers. Jun kinases such as JNK2 (MAP kinase 9) mediate signal transduction of pro-inflammatory cytokines and cellular stress (Uciechowski *et al.*, 1996).

CD40 is a receptor on the surface of B-lymphocytes, the activation of which plays critical role in B cell proliferation and differentiation. CRAF1, (CD40 receptor-associated factor 1) encodes a protein that interacts directly with CD40 receptor (Cheng *et al.*, 1995). Its upregulation in the co-culture is perhaps a main determinant of lymphoepithelial crosstalk as discussed above.

The c-myc gene is commonly amplified and over-expressed in many human tumors (Ryan and Birme, 1996). A member of the myc family of helix-loop-helix transcription factors, c-myc is integral in controlling cell growth and promotes cell proliferation and transformation by activating growth-promoting genes (Thompson, 1998). Prothymosin- α (PT- α) is a nuclear protein and its expression is associated with alterations in the proliferative state of cells and has been reported to be regulated by the c-myc gene in vitro. (Smith, 1995; Mon *et al.*, 1993). The increased activity of c-myc in this model is likely to result in the increase in RT- α mRNA.

PKC- α protein levels regulate certain pathways that lead to the expression of differentiation-dependent genes. In a series of antisense transfection experiments where PKC- α expression in CaCo-2 cells was almost completely deleted, enhanced proliferation and a marked decrease in differentiation was observed, as well as a more aggressive transformed phenotype (Scaglione-Sewell *et al.*, 1998). In a similar fashion, the lack of PKC-

α mRNA detected in the co-culture 'M cells' may underlie some of the phenotype changes featured.

5 Glutathione S-transferase A1 (GSTA1) is a member of a multigene family of detoxification and metabolizing enzymes. Induction of GST enzyme activity has been demonstrated to act as a potent anti-proliferative and differentiating agent in Caco-2 cells (Stein *et al.*, 1996) suggesting a similar role in the 'M cell.'

 Transcription factor GATA-2 is thought to maintain and promote the proliferation of early haematopoietic progenitor cells.

10 The placenta growth factor (PLGF) is a member of the vascular endothelial growth factor (VEGF) family of growth factors. In addition to PLGF, VEGF mRNA was enhanced in the co-culture cells. These growth factors play a crucial role in angiogenesis during development and/or repair (Andre *et al.*, 2000). The augmented transcription of their mRNA is consequently not a surprising find. However, hypoxia and energy depletion are known to induce angiogenesis by increasing VEGF, expression and so the possibility that the co-culture conditions are responsible for these genes induction cannot be ruled out rather than
15 a deliberate mechanism of neogenesis in M cell formation. VEGF receptor 1 (VEGFR1); the receptor for VEGF and PLGF, mRNA is down-regulated and is possibly a consequence of desensitization of the receptor by VEGF and PLGF binding, initiating a reduction in the receptor's RNA.

20 Coinciding with the above actions, the absence of growth factors such as insulin-like growth factor-binding protein 3 (IGFBP3) and keratinocyte growth factor (KGF) may be modulating enterocytic cell proliferation and differentiation.

 Caco-2 cells have been shown to express the type I IL-1R. (Varilek *et al.*, 1994) IL-1R α binds IL-1 and mediates cell signalling particularly signalling involved in cell proliferation
25 (French *et al.*, 1996). The expression of IL-1R can be enhanced by IFN- γ (Varilek *et al.*, 1994). Therefore, the expression of IL-1R type 1 mRNA in the co-culture is interesting when considering the significant expression of IFN-1 expressed in supernatants of the co-culture model.

30 In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, PKC- α , GSTA1, GATA-2, and PLGF.

Differentiation

Development of cells or differentiation is dictated by the expression of a cell's genes specific to that cell. This is a particularly important aspect with regards to M cells.

5 The cortical cytoskeleton not only provides structural support to the plasma membrane but also contributes to important dynamic processes such as endocytosis, exocytosis, and transmembrane signalling pathways. Ezrin, or villin 2, is an F-actin associated molecule and is concentrated in surface projections such as microvilli and membrane ruffles where they link the microfilaments to the membrane and has been
10 reported to be in abundance during development and differentiation of the intestinal epithelium. It was reported that hepatocyte growth factor (HGF/SF) could stimulate the tyrosine phosphorylation of ezrin in a human colon epithelial cell line, which induced the ezrin associated membrane ruffling. It is interesting to note that both hepatocyte growth factor activator (HGF activator) and hepatocyte growth factor-like protein were both upregulated in
15 the co-culture model and taken with the augmented ezrin mRNA the induction of these genes would appear to underlie the mechanism involved in the morphogenesis observed in M-cells.

 These data demonstrate that the expression of the ezrin gene is being regulated at the level of mRNA due to effects incurred by the B-cells. It is particularly relevant
20 considering the observations of villin diffusely displayed in M-cells.

 One method of actin cytoskeletal reorganization is controlled by the LIMK-1 serine/threonine kinase, which acts by phosphorylating cofilin and subsequently Rac (as previously reported). However, LIMK-1 was deleted in the co-culture model and would appear to rule out the Rac-mediated mechanism of actin reorganization in the M cell model.

25 The cadherin family of cell adhesion molecules play important role in cell-cell adhesion during tissue differentiation. They have been reported to be linked to the actin cytoskeleton by catenins located in the cytoplasmic compartment of the cell. The specific expression of NCAD in the co-culture suggests a distinct gene involved in the cytoskeletal structure.

30 Previous reports have shown that neogenin is closely related to the human tumor suppressor molecule DCC (deleted in colorectal cancer) and together they constitute a subgroup of Ig superfamily proteins that have shown to be essential for terminal differentiation of specific cell types in the adult including the human colon. These parallels suggest that neogenin, like DCC, is functionally involved in the transition from cell

proliferation to terminal differentiation of specific cell types. Its absence in the co-culture model might represent a period of continued proliferation for the cells and allow a longer period of proliferation.

The helix-loop-helix (HLH) family of transcription factors has been shown by others to play a central role in the regulation of cell growth, differentiation and tumorigenesis. Of particular interest, when HLH 1R21 was over-expressed in mouse NIH3T3 cells, it induced a morphologically transformed phenotype.

Other genes associated with differentiation including Myeloid cell nuclear differentiation antigen (MNDA) and the LHX1 gene. The LHX1 gene is a member of the LIM/homeobox (Lhx) gene family. It has been shown that it codes for a transcriptional regulatory protein involved in the control of differentiation and development.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of HGF activator, ezrin, NCAD, MNDA, and LHX1.

Adhesion

It is clearly evident that modification of the M cell apical surface is a determining factor in M cell apical membrane adherence, and thus, uptake and transport of macromolecules/microorganisms and targeting epitopes on the surface of M cells has been used to promote further adherence and uptake of particles in vaccinology. The specificity of these markers is not only useful for vaccine strategies but also represents targets for understanding adhesion and uptake of bacteria and viruses. Adhesion is not privy to the apical surface. Adhesion molecules on the basolateral surface of M cells, such as cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4 are understood to be involved in leukocyte migration and in the development/organization of lymphoid nodules in Peyer's patches. Genes expressed/induced in the co-culture can provide an insight into the mechanisms involved and are discussed below.

The tyrosine kinase receptor TIE 1 is normally located in vascular endothelial and haematopoietic cells and is largely involved in the proliferation and differentiation of miniature haematopoietic cells and would be an appropriate gene specific for M cells. In the brain, TIE

mRNA and protein is significantly elevated in lesions composed of abnormal vasculature called arteriovenous malformations (AVMs) and the surrounding vasculature. Like AVMs, the significant upregulation of TIE in M cells may indicate some ongoing neogenesis, and depending on the receptor's polarity could be of potential use in vaccine targeting.

5 The neuronal cell adhesion molecule L1 (NCAML1) is a transmembrane glycoprotein belonging to the immunoglobulin superfamily and is generally associated with development of the nervous system. As a potent promoter of neurite growth, it is allied with plastic changes. In nerve growth it interacts with the actin cytoskeleton via an ankyrin linkage and promotes specific distribution of F-actin. Such flexibility is ideal in the M cell scenario.

10 The integrin family consists of a series of related alpha beta heterodimers involved in a variety of cell-matrix and cell-cell adhesion functions. The $\alpha_3\beta_1$ integrin is a multiligand extracellular matrix receptor found on many cell types and can function as a receptor for fibronectin, laminin, and collagen. Phagocytosis of molecules by breast cells has also been reported to involve this adhesion molecule, thus, it would appear a suitable candidate as an

15 adhesion target on M cells.

 The leukocyte adhesion glycoprotein p150 (CD11C antigen), also a member of the integrin family, is involved in leukocyte sequestration via interaction of CD11/CD18 similar to that of ICAM-1.

20 In stratified epithelia β_4 integrin (CD104 antigen) has been shown to be important for proper differential expression and crucial for stable adhesion to the basement membrane through its ability to attach externally to laminin and internally to the keratin cytoskeleton. Interestingly, during human intestinal organogenesis receptors have been shown to occur. This integrin would appear to play an important role in epithelial cell-matrix interactions during development but particularly in M cell development.

25 CD44 is a major surface adhesion molecule involved in cell-cell and cell-matrix interactions and lymphocyte homing and activation. The observed enhanced expression suggests that this molecule is an important feature in the activities of M cells. A non-receptor tyrosine kinase, C-src protooncogene (SRC1) has been shown to cause overexpression of CD44 in the intestine. As well as its effects on proliferation, the enhanced

30 activity of SRC1 seen in the M cell model would appear to have major effects on cell adhesion properties of the M cell. Hepatocyte growth factor activator (HGF activator) is a serine protease produced and normally secreted by the liver. It has been documented as stimulating reparative processes in intestinal epithelial cells and could be why its activity is enhanced in this model. However, stimulation of CD44 in colonic epithelial cells has been

reported to augment c-met, the HGF receptor. This in turn stimulates the "inside-out signalling causing an amplified expression of integrins that leads to an increase in vascular adhesion to the epithelium.

5 It has been reported that the glutamate receptor (NMDA) is generally associated with learning and memory, highly plastic processes in the brain. The high density of NMDA receptors reflects similar plastic changes seen in the co-culture model but would also act as a target epitope for drug delivery.

10 TKT is a tyrosine-kinase receptor related to TRK and is a member of cell adhesion kinase receptor family. Ephrin (type A) is a tyrosine kinase receptor that has been reported to be involved in neogenesis and tumor formation. Sp1 is a nuclear protein constitutively expressed and mediates basal promoter activity and is the main Vitamin-D receptor promoter in intestine. These are all potential target sites relevant to M cells.

15 Many of the receptors/cell surface antigen 'deleted' (not detectable) in the co-cultures could be putative negative markers of M cells. A good example is the laminin receptor $\alpha_7\beta_1$ integrin. Expression of the $\alpha_7\beta_1$ integrin correlates with human intestinal cell differentiation and could be used in a similar fashion that was applied with sucrase isomaltase and alkaline phosphatase.

20 In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAM1, $\alpha_3\beta_1$ integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1.

25 Transport

30 The RAB proteins are reported to be regulators of polarized membrane traffic in epithelial cells. The RAB3B is localized to the apical pole very near the tight junctions between adjacent epithelial cells where it is reported to be a possible regulator of apical and/or junctional protein traffic in epithelial tissues. RAB3B is highly homologous to a brain-specific RAB3 isoform (RAB3A) that targets the presynaptic nerve terminal, where it is reported to regulate exocytosis.

In polarized cells, the small GTPase Rab5a is localized to the plasma membrane, clathrin-coated vesicles, and early endosomes and is a regulator of transport between the

plasma membrane and early endosomes. The decreased expression of RAB5a seen in the co-culture may deregulate the rate of endocytosis and/or vesicle fusion and could possibly release 'the brake' on vesicle trafficking.

5 RAB6 is another ras related protein also a regulator of intracellular transport in mammalian cells. It controls intra-Golgi transport, either acting as an inhibitor in anterograde transport or as a positive regulator of retrograde transport. Like RAB5a, the pronounced decrease seen in mRNA transcription could be a means of subverting transport regulation in epithelial cells and so optimize the process as observed in M cells.

10 Protein kinase C (PKC) and the actin cytoskeleton are critical effectors of membrane trafficking in mammalian cells. The F-actin cross-linking protein myristoylated alanine-rich C kinase substrate (MARCKS), a substrate for PKC, has been reported to be a component of the mechanism of endocytosis.

15 TIR or p71 plays a key role in the control of cell proliferation through the binding of transferrin, the major iron-carrier protein. Located on both apical and basolateral surfaces, the transferring receptor has the ability to internalize and recycle to the surface. Indeed experiments by Hughson and Hopkins (1990) demonstrate pathways from the apical and basolateral surfaces meet in an endosomal compartment. Furthermore, Shah and Shen (1994) discovered that the fungal metabolite brefeldin A (BFA) could relocate receptor distribution and enhance TfR mediated transcytosis. The increased expression of this
20 mRNA in the M cell model suggests a potential delivery mechanism of protein drugs across the intestinal epithelium present in M cells that could be exploited.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of a RAB protein, PKC, and TfR.

25

Signal transduction

30 In order for a cell to respond to extracellular signals, which cause it to alter gene expression or cellular function, it must involve the activation of a signal transduction cascade. There are many different types of signalling cascades, which can be unique to a specific type of stimulus. There are two main mechanisms by which these cascades transmit their signal, either through the regulation of enzymes, which produce second messenger molecules or through the regulation of protein phosphorylation. The activation of these cascades is usually mediated through specific cell surface or intracellular receptor proteins. The receptor

protein recognizes the incoming extracellular signal and responds accordingly, initiating a specific series of intracellular signal that direct the cell's behavior. A number of genes involved in intracellular signalling were upregulated or induced in the M cell model and are discussed below.

5 A member of the Janus family of tyrosine kinases, which are non-receptor protein kinases, Jak3 is involved in intracellular signalling mediated by cytokines and growth factors such as IL-2, IL-4, and IL-7. Jak3 has been reported to play a crucial role in Peyer's patch organogenesis. Mutant mice deficient in Jak3 presented defects in lymphocyte production and the absence of Peyer's patch structures. Its induced expression suggests a greater
10 level of activity and possibly a major requirement underlying the M cell phenotype 'switch'.

 The nuclear zinc-finger transcription factor, early growth response factor-1 (EGR-1) is an immediate-early gene product expressed in response to diverse stimuli and is involved in growth, development, and differentiation. EGR-1 has been reported to function in growth regulation and suppression of cell transformation by transactivation of the TGF β gene.
15 TGF β is capable of stimulating the synthesis of extracellular matrix proteins that can potentially stabilize epithelial cell contact with the substratum. In addition EGR-1 also plays a role in the immune response, regulating targets such as IL-2, CD44, ICAM-1, and TNF. Taken together the considerable induction of EGR-1 mRNA emphasizes the importance of this protein's involvement in M cell behavior.

20 CaM kinase IV (CAMK IV) is involved in Ca²⁺-dependent mechanism for regulating MAP kinase pathways. Many kinases activity has been observed to be enhanced in this model and so it is logical that CAMK IV expression is induced as a requirement to function.

 The tyrosine kinase Trk1 has been reported to be involved in signalling pathways involving development in adult tissues and in cells of the lymphohaematopoietic system.

25 Epithelial discoidin domain receptor 1 (EDDR1) mRNA was reduced in the co-culture. EDDR1 is a collagen receptor involved in controlling cellular responses to the extracellular matrix (ECM). The decrease in this gene would implicate it in the reorganization of the M cell in relation to the ECM.

 cAMP-dependent protein kinase type I beta regulatory subunit (PRKAR1B) stimulates
30 growth by modulating the signalling of camp via its regulation of cAMP-dependent protein kinase (PKA). PRKAR1B's reduction in the co-culture model may represent an inhibitory role in the cell's growth counterbalancing the proliferative signals.

 In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the

protein may be selected from the group consisting of Jak 3, EGR-1, TNK1 and CAMK IV.

Protection and repair

5 Chaperones such as HSP40 and HSP 70 participate in many biological processes in which protein folding is involved. These include protein translocation, protein translation, protein assembly and disassembly, and protein degradation. It is understandable that such genes would be induced considering the evolving processes of a phenotype 'switch.' However heat shock protein production has been reported to be induced as a result of harsh
10 changes in their environmental conditions such as stress, ischaemia or hypoxia resulting in protein damage. Therefore it cannot be ruled out that the induction of these genes is in fact a protective measure as a consequence of the adverse conditions in the co-culture.

 HSP 60 has been observed in highly replicating cells e.g. short-living epithelial cells of the intestine. Involved in the import and refolding of nuclear-encoded proteins destined for
15 the mitochondrial matrix.

 The 27-kDa heat shock protein (HSP27) is expressed in a variety of tissues, including gut epithelia and in the absence of stress has been reported to regulate actin filament dynamics. Hsp27 induction in the M cell model like the other heat shock proteins (HSPs) may be active in development of resistance to stressful conditions. Activation of HSP27 can
20 contribute to agonist-induced phosphorylation-modulated reorganization of the actin cytoskeleton and, in the case of stress activation, provides an actin-based adaptive response of cells to the new environmental conditions, and is ideal candidate for the plasticity seen in M cells.

 Expression of receptors for fMLP on human phagocytes is well established, but there
25 is conflicting evidence regarding the potential expression of fMLP receptors on other cells within the mucosa, particularly the epithelial cells. The reported observation of the receptor for the chemotactic peptide fMLP supports the notion of the intestinal epithelial cell as an early "sensor" of infection and inflammation. It has been reported that, fMLP, present in abundance in the lumen of the gut and that activation of fMLP receptors induces cytotoxic
30 effects such as lysosomal release and superoxide generation. Thus, it would appear that their presence would be a defensive role in the event of infection of microorganisms.

 Glutaredoxin (thioltransferase) is a small, heat-stable protein catalyzing glutathione-dependent disulfide oxidoreduction reactions in a coupled system with NADPH, GSH and glutathione reductase. It is important in regulating cell metabolism through the inactivation of

oxidated transcription factors thought to be important in cellular responses to oxidant stress. This modulation of transcription factors' binding activity has been demonstrated for a number of transcription factors, including NF-kB/Rel proteins, Fos and Jun proteins and nuclear factor I (NFI) family of transcription factors. The induction of such a gene would appear to provide a protective role and is particularly influential on a number of key transcription factors.

CREB has been implicated as having prominent role in protection. Over-expression of the gene was reported to reverse hypoxia elicited TNF induction. This infers that the increase in the cAMP responsive element binding protein (CREB) mRNA is possibly a protective response to conditions.

Inactive in cells under normal conditions, gadd153 expression is markedly induced in response to a variety of cellular stresses, including nutrient deprivation, DNA damage, and oxidative stress (e.g. free radicals) which normally leads to growth arrest. The arrest in growth is thought to allow critical repair processes to be carried out before any further cell cycling. It would appear that the gadd153 expression in the co-culture is for reparative purposes.

The excision repair proteins XPG and XPD have been reported to be involved in nucleotide repair. In addition, mRNA for ubiquitin-conjugating enzyme (likely to be involved in post-replication repair and induced mutagenesis, RAD23, and ataxia telangiectasia are also expressed in the co-culture. Their expression, coinciding with gadd153 suggests there is a high degree of impairment to genes in the M cell model.

Interleukin-13 (IL-13) is a potent anti-inflammatory cytokine and has been reported to have the same protective properties in inflammation as IL-4 through its ability to modulate and suppress pro-inflammatory cytokines. It is puzzling that in an environment with a high level of pro-inflammatory cytokines produced that IL-13 mRNA is in fact reduced. One possible explanation might be its anti-adhesion effect. It has been reported that IL-13 (secreted from lymphocytes) down regulated cell adhesion molecules in colonic epithelium and so the role of IL-13 in the co-cultured cells is modulating cell adhesion properties and not inflammation.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd 153, XPG, XPD, ubiquitin, conjugating enzyme, RAD 23, and ataxia telangiectasia.

Apoptosis and programmed cell death

In programmed cell death, apoptosis is programmed in the sense that a genetically directed 'clock' selects a given time for the death of certain cells. It has been reported that it provides an important mechanism for the maintenance and renewal of cells in the gut and in development. However, for the epithelium to maintain its barrier functions, the level of apoptosis needs to be regulated, and this is 'checked' by several signal transduction systems. Toxic insult or lack of factors that maintain cell survival can also lead to apoptotic death of the cell.

It has been reported that over-expression of c-fos and c-jun (constituents of the AP-1 transcription factor) in the intestine correlates with programmed cell death and subsequent cellular regeneration. Other studies have demonstrated increases in both proximal jejunum and colon jun mRNA level coincide with a period of major changes in intestinal cell proliferation). The c-jun protein product involved in activation of AP-1, transcription is enhanced when it is phosphorylated by stress-activated protein kinases of which there are many in the M cell model.

As intestinal epithelial cells reach the villus apex they undergo apoptosis and, are shed and, in normal circumstances, caspases, a family of cysteine proteases, play a central role in initiating, amplifying, and executing apoptosis. The pattern of caspase activation in this process is not understood. It is interesting to note that the apoptosis regulator, bcl-x, and caspase 9 are induced in the co-culture. The bcl-x gene plays an important role in the regulation of programmed cell death (PCD), depending on its splice variant the bcl-x protein can accelerate apoptosis or delay/prevent programmed cell death (as previously reported). Bcl-x controls apoptosis mechanisms at points upstream of caspase activation. Perhaps, it is responsible for the marked induction of caspase-9. Caspase-9 is a caspase initiator. Once activated, it can proteolytically activate other caspases (including 3, 6 and 7), which in turn activate caspase-2 and 6 (as previously reported). Inhibitor of apoptosis protein 2 (IAP2) binds to and inhibits caspase-3. Its expression is a mechanism of regulating cell death depending on the particular cellular or environmental signals. Therefore, its absence in the co-culture cells and the increased activity of caspase-9 allows caspase-3 unchecked pro-apoptotic activity.

The death domain receptor 3 (DDR3) member of the TNFR family can induce apoptosis as previously reported. Its mRNA expression is also reduced in the co-culture model.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group selected from: bcl-x and capase-9 and more generally in view of the foregoing may be selected from the group consisting of

5 cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, CD40, C-MYC, PKC- α , GSTA1, GATA-2, PLGF, ezrin, HGF activator, hepatocyte growth factor-like protein, NCAD, MNDA, LHX1, TIE-1, NCAML1, CD104, CD44, SRC1, NMDA, TKT, ephrin (type A), Sp1, RAB proteins, PKC, TIR, Jak3, EGR-1, TNK1, CAMK IV, HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin,
10 CREB, gadd153, XPG, XPD, ubiquitin- conjugating enzyme, RAD23, cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, $\alpha\beta$ 1 integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1, a RAB protein, PKC, and TfR, bcl-x and capase-9

Example 6

15

Targeted Gene delivery

Delivery of genes, gene fragments, oligonucleotides or other nucleotide fragments or analogues of the present invention to a living organism can be accomplished by methods currently available in the prior art. For example, various recombinant viruses have been
20 used for the oral delivery of genes, such as adenovirus, retrovirus, adeno-associated virus, vaccinia virus, lenti-virus and plant-derived viruses, wherein the viral genome is replaced with an expression vector for the gene of interest. . See, David T. Page and Sally Cudmore (2001). Innovations in oral gene delivery: challenges and potentials. *Drug Discovery Today*, Vol. 6, No. 2, pp 92–101. Viral mimetic particles such as virosomes and various types of
25 polymers and liposomes, such as cationic and fusogenic, are also employed for gene delivery. See, U.S. Patent Nos. 4885172, 5047245, 5171578, 5059421, 5399331, 5204112, 1252263, 5376452, 5552155, 6120797, 6087325, 6143716. Examples of polymers are PLGA, PLA co-polymers, chitosan, and fumaric acid/sebacic acid co-polymers. For these systems, the polymer or liposome is formed from component parts in a solution of the gene
30 expression vector, thus encapsulating the genes when particles are formed. Cationic lipids such as DOTAP and polyethylenimine are commonly used whereby the gene expression vector is complexed with and protected by the lipids. (See, Ogris M. *et al.* (2001). DNA/polyethylenimine transfection particles: Influence of ligands, polymer size, and pegylation on internalization and gene expression. *AAPS PharmSci*, 3 (3), article 21).

Agents such as protamine are used to condense DNA, which due to the reduction in size of the DNA particles are more easily taken up by cells. Recombinant live bacteria (e.g. *Shigella* spp., *Salmonella* spp.) have also been exploited for gene delivery to the gut. Oral bioavailability enhancers, (e.g. sodium caprate, Elan's PROMDAS technology) could be used to increase uptake of a gene or encapsulated gene formulation.

In all cases the delivery systems can be targeted with various ligands on the surface of the particles in order to enhance binding to specific cells type and/or to enhance uptake. These ligands could be peptides, proteins, antibodies, peptidomimetics, and lipids that recognize or are being recognized by specific sites/receptors on the cell surface (Maruyama K. (2000). *In vivo* Targeting by Liposomes. *Biol. Pharm. Bull.*, 23(7), 791-799).

The targeting ligands may be peptide based, peptidomimetic based, antibody based, single chain antibody based, small organic molecule based. The targeting ligands may also be natural substrates for such receptors, transporters or other cell surface molecules found on the surface of M cells or other cell types found in Peyer's patch. The targeting ligands may be engineered so as to be genetically expressed on the surface of viruses, bacteriophages, virosomes, bacteria or other organisms, which can be utilized for vaccine delivery in the gut. Furthermore the targeting ligands can be presented either as direct conjugates to antigens, or on the surface of drug-loaded particulates such as liposomes, PLGA particles, other particulates and at the same time retain recognition by and interaction with the receptors, transporters or other cell surface molecules found on the surface of M cells and / or other cells of Peyer's patch tissue.

Examples of peptides that target the gastro-intestinal tract, in particular, membrane translocating peptides useful for vaccine delivery to M cells along with M cell specific targeting ligands are described in Table 4.

Further, targeting ligands can be genetically engineered into the surface coats of viruses, bacteriophages or bacteria, conjugated directly to antigens conjugated to the lipids in liposomes by covalent methods or streptavidin-biotin linkages, or coated onto the surface of polymers after particle formation (Torchilin V.P. *et al.* (2001) *Proc. Natl. Acad. Sci. USA*, Vol. 98, Issue 15, 8786-8791, July 17).

TAT peptide on the surface of the liposomes affords their efficient delivery even at low temperature and in the presence of metabolic inhibitors; Lestini *et al.* (2002). Surface modification of liposomes for selective cell targeting in cardiovascular drug delivery. *J. Controlled Release*, 78, 235-247; Dokka S. *et al.* (1997) Cellular delivery of oligonucleotides by synthetic import peptide carrier. *Pharm. Res.*, vol. 14, No. 12, 1759-1764; Wu Y *et al.*

(2000). Gene transfer facilitated by a cellular targeting molecule, retrovirus protein α 1. *Gene Therapy*, 7, 61-69).

5 When the delivery of the gene to M cells in the gut is designed to prime or boost the immune system, the genes can be co-delivered/co-encapsulated with adjuvants (e.g. MF59, alum, saponin, QS21, MPL, bacterial toxins such as Lt, CT or mutants there-of, CPG motif nucleotides). Immune response could be boosted at a later stage by methods such as subcutaneous administration of an adjuvant.

10 In some cases it may be desired to shut off expression of certain genes, so as to enhance the adoption by enterocytes of an M cell phenotype. This can be achieved by the delivery, by methods outlined above, of antisense oligonucleotides, ribozyme, or RNA-interference molecules specific to the gene of interest.

Table 4
Peptides that Target to and/or Enhance Uptake Across the GIT

SEQ. ID. NO:	PEPTIDE SEQUENCES
SEQ. ID NO:	ADDFMGCMLTLPTSLGGEGSPYNYDTHEANGPH
SEQ. ID NO:	TPTTTATVVGTTGPVDLSSLHLLRHPCREF
SEQ. ID NO:	MSPDHQYALQSSPVLPCCRPLLVDSDYIHS
SEQ. ID NO:	RGYGRLAESCCVNCIRTVGGCGNSPASDILSAT
SEQ. ID NO:	STPGRGSGRDTGANNPADTPYANPSHRDTILSLDPSLL
SEQ. ID NO:	RQHLVVRDLHGPRFRDNTGAGTFSPPVSVADTHRTPD
SEQ. ID NO:	SFSNLTAGDEEDDHFSGGRFNHNALTSRSHNRGQLASSA
SEQ. ID NO:	RQSVLDSWGGKTSVTGLSERYYASHSHTSAPTPHYASHS
SEQ. ID NO:	RQWVGDRDAGEGNTWVDEKYSRDANVISYRSHNHASQGT
SEQ. ID NO:	RASDCDVECNLRWVEDVGGVWYAKTVSRMLSTT
SEQ. ID NO:	RQSAGFLGFAPTNIIDTSFNAGCGDTLAIPCRHRSSLISPARPP
SEQ. ID NO:	RSGAYESPDGRGGRSYVGGGGCGNIGRKHNWLWGLRTASPACWD
SEQ. ID NO:	SPRSFWPVVSRHESFGISNYLGCGRYTCISGTMTKSSPIYPRHS
SEQ. ID NO:	SSSSDWGGVPGKVVRERFKGRGCGISITSVLTGKPNPCPEPKAA
SEQ. ID NO:	RVGQCTDSDVRRPWARSCAHQCGGAGTRNSHGCITRPLRQASAH
SEQ. ID NO:	SHSGGMNRAYGDVRELDRDRWNATSHHTRPTPQLPRGPN
SEQ. ID NO:	SPCGGSWGRFMQGGFLGGRTDGCGAHRNRTSASLEPPSSDY
SEQ. ID NO:	RGAADQRRGWSNGLPRVGDWDAIAHNSYFTTSRRPRPP
SEQ. ID NO:	SGGEVSSWGRVNDLCARVSWTGCGTARSARTDNKGFLPKHSSLR
SEQ. ID NO:	SDSDGDHYGLRGGRVRCSLDRGCGLALSTVHAGPPSFYPKLSSP
SEQ. ID NO:	RSLGNYGVTGTVDVTLPMPGHANHLGVSSASSSDPPRR
SEQ. ID NO:	RTTTAKGCLLGSFGVLSGCSFTPTSPPHLGYPPHSVN
SEQ. ID NO:	SPKLSSVGMVKTELPTGPNNAISIPISATLGPRNPLR
SEQ. ID NO:	RWCGAELCNSVTKKFRPGWRDHANPSTHHRTPPPSQSSP
SEQ. ID NO:	RWCGADDPCGASRWGRGNSLFGCGLRCSAAQSTPSGRIHSTSTS
SEQ. ID NO:	SKSGEGGDSSRGETGWARVRSHAMTAGRFRWYNQLPSDR
SEQ. ID NO:	RSSANNCEWKSOWMRRRA,CIARYANSSGPARAVDTKAAP
SEQ. ID NO:	SKWSWSSRWGSPQDKVEKTRAGCGGSPSTNCHPYTFAPPPQAG
SEQ. ID NO:	SGFWEFSRGLWDGENRKSVRSGCGFRGSSAQGPCVTPATIDKH
SEQ. ID NO:	SESGRCRSVSRWMTTWQTQKGGCGSNVSRGSPLDPSHQTGHATT
SEQ. ID NO:	REWRFAGPPLDLWAGPSLPSFNASSHPRALRTYWSQRPR
SEQ. ID NO:	RMEDIKNSGWRDSCRWGDLRPGCGSRQWYPSNMRSSRDYPAGGH
SEQ. ID NO:	SHPWYRHWNHGDFSGSGQSRHTPPESPHPGRPNATI
SEQ. ID NO:	RYKHDIGCDAGVDKSSSVRGCGAHSSPPRAGRGRGTMTVSRL
SEQ. ID NO:	SQGSKQCMQYRTGRLTVGSEYCGGMNPARHATPAYPARLLPRYR
SEQ. ID NO:	SGRTTSEISGLWGWGDDRSYGWGNLTPNYIPYRQATNRHRYT
SEQ. ID NO:	RWNWTVLPATGGHYWTRSTDYHAINNHRPSIPHQHPTPI
SEQ. ID NO:	SWSSWNWSSKTTRLGDRATREGCGPSQSDGCPYNGRLTTVKPRT
SEQ. ID NO:	SGSLNAWQPRSWVGGAFRSHANNLNPKPTMVTTRHPT
SEQ. ID NO:	RYSGLSPRDNGPACSQEATLEGCGAQRLMSTRRKGRNSRPGWTL
SEQ. ID NO:	SVGNDKTSRPVSFYGRVSDLWNASLMPKRTPPSKRHDDG
SEQ. ID NO:	TNAKHSSHNRRLRTR
SEQ. ID NO:	SDNAKEPGDYNCCGNGNSTG
SEQ. ID NO:	RTRLRRNHSSHKANT

SEQ. ID. NO.:	PEPTIDE SEQUENCES
SEQ. ID NO:	GPHRRGRPNRRSSKT
SEQ. ID NO	GTSNGNGCCNYDGP
	<u>Peyers patch and/or M cell specific targeting ligands:</u>
SEQ. ID NO:	ATPPPWLLRTAP
SEQ. ID NO:	DGSIHKRNIMPL
SEQ. ID NO:	DYDSLWRSTLH
SEQ. ID NO:	GEPTTDMRWRNP
SEQ. ID NO:	GLWPWNPVTVLP
SEQ. ID NO:	HMLNDPTPPPYW
SEQ. ID NO:	KPAYTHEYRWLA
SEQ. ID NO:	LETTCASLCYPS
SEQ. ID NO:	LGTDWHSVSYTL
SEQ. ID NO:	LGTNAGVPGFP
SEQ. ID NO:	LTHSKNPVFLST
SEQ. ID NO:	LVPTTHRHWPVT
SEQ. ID NO:	LVSNARGFNNLS
SEQ. ID NO:	NTRIEPIRFYM
SEQ. ID NO:	NVYTFHSMSPMP
SEQ. ID NO:	QHTTLTSHPRQY
SEQ. ID NO:	SDFSDTMPHRPS
SEQ. ID NO:	SIDTIQILSLRS
SEQ. ID NO:	SISWASQPPYSL
SEQ. ID NO:	SMVKFPRPLDSR
SEQ. ID NO:	SPTLGASVAQTN
SEQ. ID NO:	TMSPNVYYTAFG
SEQ. ID NO:	TQIPSRPQTSPQ
SEQ. ID NO:	VCSNMYFSCRLS
SEQ. ID NO:	VPPHPMTYSCQY
SEQ. ID NO:	VPRLEATMVPDI
SEQ. ID NO:	VPTKPELPVNFT
SEQ. ID NO:	WSSDLPQPASTY
SEQ. ID NO:	YITPYAHLRGGN
SEQ. ID NO:	NVYTDNTLSPTP
SEQ. ID NO:	LETTAASLCYPS
SEQ. ID NO:	SPYCLSACTTEL
SEQ. ID NO:	LETTCASLCYPS
SEQ. ID NO:	VPPHPMTYSCQY
SEQ. ID NO:	VPPHPMTYSAQY
SEQ. ID NO:	VPPHPMTYSSQY
SEQ. ID NO:	YQCSYTMPPPV
SEQ. ID NO:	VCSNMYFSCRLS
SEQ. ID NO:	VSSNMYFSSRLS
SEQ. ID NO:	DYDSLWRSTLHGGHESH
SEQ. ID NO:	GNPTSTMRAW
SEQ. ID NO:	PWNSATVL
SEQ. ID NO:	NDPTAPPY

SEQ. ID. NO:	PEPTIDE SEQUENCES
	<u>Membrane Translocating Peptides:</u> (underline denotes cyclization)
SEQ. ID NO:	KKAAAVLLPVLLAAP FITC-LC
SEQ. ID NO:	KKKAAAVLLPVLLAAP
SEQ. ID NO:	KKAAAVLLPVLLAAPREDL
SEQ. ID NO:	KKCAAVLLPVLLAAPC
SEQ. ID NO:	CAAVLLPVLLAAC
SEQ. ID NO:	KKCAAVLLPVLLAC
SEQ. ID NO:	CAAVLLPVLLC
SEQ. ID NO:	CAAVLLPVLC
SEQ. ID NO:	CAVLLPVLLAAPC
SEQ. ID NO:	CVLLPVLLAAPC
SEQ. ID NO:	CLLPVLLAAPC
SEQ. ID NO:	CLPVLLAAPC
SEQ. ID NO:	AAVLLPVLLAAP
SEQ. ID NO:	AAVLLPVLLAA
SEQ. ID NO:	KKAAVLLPVLLA
SEQ. ID NO:	AAVLLPVLL
SEQ. ID NO:	AAVLLPVLL
SEQ. ID NO:	AVLLPVLLAAP
SEQ. ID NO:	VLLPVLLAAP
SEQ. ID NO:	LLPVLLAAP
SEQ. ID NO:	LPVLLAAP
SEQ. ID NO:	AAVLLPVLLAAKKRKA
SEQ. ID NO:	KKKRKAAAAPVLLA

Example 7

5

Use of bacterial coatings to convert enterocytes to M cells

Might be nice to have some type of claim capturing the concept from this section Use of bacterial coatings on PLGA particles, co-administered bacterial particles or pro-biotic yogurts as adjuvants for oral vaccination with PLGA particles. The invention is based on converting enterocytes to M cells by using specific bacteria in advance of, or along with the oral vaccine particle of interest. In doing so the capability of absorbing particles through M cells will be increased. This idea is not based on targeting but on the ability of live bacteria or active bacterial components to stimulate cytokine production in Peyer's patches, thus, enabling enterocyte-M cell conversion. As a result, an invention disclosed herein is a method of promoting enterocyte-M cell conversion, said method comprising orally administering an antigen, antigenic composition, or antigen-carrying particle to a person and either simultaneously with, or prior to, said administration, also orally administering a

15

bacteria, or pro-biotic yogurts, or bacterial component to said person.

All references cited herein are incorporated herein by reference in their entireties.

5

Table 5
Miscellaneous GenBank Accession Numbers

Human Serum Albumin	NM_000477.3
Calreticulin	M84739

10

15

Dates for GenBank records

To the extent the date of a GenBank record, rather than its version number, is relevant for purposes of incorporation by reference, the date of the record is the filing date of this application with the following exceptions:

20

Table 2: Rat genes

3/27/02	D83697 through M10149
3/28/02	Q03238 through NM_017218.2

25

Tables 3: Human genes with a fold change of 0.5 or less
4/02/02

Table 2 Human genes with a fold change of 0.5 or less

4/02/02	U76376.1 through XM_087242.1
4/03/02	S90469 through M29366.1

30

The records specified for 3/27/02, 3/28/02, 4/02/02, and 4/03/02, do not include those

of GenBank IDs: Q07912, P21145, P46734, Q92851, Q13490, NP_006168, P28360, P28347, Q06830, P20701, P01589, P05106, P35225, P17936, S18408, P17074, P10661, P35426, P09456, P22791, Q06486, P21708, Q03238, P10644, P54868, P10398, P48730, P31749, P27361, P25063, Q14012, P13866

5

CLAIMS

1. A method of increasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell a nucleic acid coding for a protein, wherein absent
5 said increase, the levels of said protein or its mRNA is greater than in a non-Peyer's patch cell.
2. The method of Claim 1 wherein the protein is a transcription factor or a protein that activates a transcription factor.
3. The method of Claim 2 wherein the transcription factor or a protein that
10 activates a transcription factor is selected from the group consisting of Jun-B; c-jun related TF, Jun-D; c-jun related TF, STAT 3 - signal transducer and activator of transcription 3, NF-kappa β Tf p105 subunit, S-myc proto-oncogene; myc related, Nm23-M2; nucleoside diphosphate kinase B; metastasis reducing protein, and C-est-I proto-oncogene; p54.
4. The method of Claim 1 wherein the protein is a receptor, or cell surface
15 antigen,
5. The method of Claim 4 wherein the protein is a receptor or a transporter.
6. The method of Claim 1 wherein the protein is selected from the group consisting of nucleoside diphosphate kinases and member of the 14-3-3 family.
7. The method of Claim 1 wherein the protein is coded for by a gene with an
20 expression Fold Change denoted by a **, *, or number greater than 2.00 in Tables 2 or 3.
8. The method of Claim 1 wherein the nucleic acid coding for at least 2 proteins is delivered, each of said proteins coded for by a gene with an expression Fold Change denoted by a **, *, or number greater than 2.00 in Tables 2 or 3.
9. The method of Claim 1 wherein the cell to which the nucleic acid is delivered
25 is a human cell.
10. The method of Claim 9 wherein the cell is in a Peyer's patch in a human and the nucleic acid is delivered by the oral route.
11. The method of Claim 9 wherein the cell is not within the body of a human.
12. The method of Claim 1 wherein the cell to which the nucleic acid is delivered
30 is a rat cell.
13. The method of Claim 1 wherein a nucleic acid coding for a tumor antigen or foreign peptide is also delivered to the Peyer's patch cell.
14. The method of Claim 13 wherein the cell to which the nucleic acid is delivered is a human cell.

15. A method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule (RNAi), said anti-sense molecule, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for said protein, wherein absent said anti-sense molecule, ribozyme or RNAi nucleic acid, the levels of said protein or its mRNA is less than in a non-Peyer's patch cell.

16. The method of Claim 15 wherein the anti-sense nucleic acid, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule is complementary to a sequence of at least 15 nucleotides of the mRNA of the protein.

17. The method of Claim 16 wherein the anti-sense nucleic acid, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule is complementary to a sequence of at least 30 nucleotides of the mRNA of the protein.

18. The method of Claim 15 wherein the protein is coded for by a gene with an expression Fold Change denoted by a "-", or a number less than 0.5 in Tables 2 or 3.

19. The method of Claim 15 comprising delivering to said cell anti-sense nucleic acid molecules, ribozyme nucleic acid molecules, RNA interference nucleic acid molecules, said anti-sense, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for at least 5 different protein a, wherein absent said anti-sense, ribozyme or RNAi nucleic acid molecule, the levels of each of said proteins or its mRNA is less than in a non-Peyer's patch cell.

20. A method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule said anti-sense, ribozyme or RNAi nucleic acid forming a double-stranded molecule with part or all of the mRNA for said protein, wherein absent said anti-sense, ribozyme or RNAi nucleic acid molecule, the levels of said protein or its mRNA is less than in a non-Peyer's patch cell.

21. A method of Claims 1, 13, or 15 in which the Peyer's patch cell is an M cell.

22. A human cell to which the method of Claims 1 has been applied, or the progeny of said human cell.

23. A human cell to which the method of Claim 13 has been applied, or the progeny of said human cell.

24. A human cell to which the method of Claim 15 has been applied, or the

progeny of said human cell.

25. A human cell to which the method of Claims 1 has been applied, or the progeny of said human cell.

5 26. A human cell to which the method of Claim 13 has been applied, or the progeny of said human cell.

27. A human cell to which the method of Claim 15 has been applied, or the progeny of said human cell.

10 28. A method for enhancing transport of a drug through the gastrointestinal tract, said method comprising orally administering said drug in a composition that comprises a transport-enhancing protein, said transport-enhancing protein selected from the group consisting of human serum albumin (HSA), clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, and Ca²⁺pla2, or a homolog that has at least 80% amino acid identity with said transport-enhancing protein over a length of said transport-enhancing protein identical to the homolog.

15 29. A method of Claim 28 wherein the homolog has at least 90% amino acid with the transport-enhancing protein over a length of the transport-enhancing protein identical to the homolog.

20 30. A method of Claim 28 wherein the transport-enhancing protein is selected from the group consisting of human serum albumin (HSA), clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, and Ca²⁺pla2.

31. A method to facilitate intracellular trafficking of an antigen that has been orally delivered by itself or as part of a composition or particle, said method comprising administering a protein selected from the group consisting of calreticulin, rab family proteins and ribosomal proteins.

25 32. A chimeric protein comprising the amino acid sequence for calreticulin, rab family proteins and ribosomal proteins and the amino acid sequence for a second polypeptide.

33. A method of administering a polypeptide, where said polypeptide is part of a chimeric protein of Claim 32, and wherein said chimeric protein is orally administered.

30 34. A method of delivering a vaccine to a target cell, said method comprising utilizing as the target cell a Peyer's patch cell in which a normally upregulated protein or mRNA is further upregulated.

35. A method of Claim 34 wherein the Peyer's patch cell is an M Cell.

36. A method of Claim 1 wherein the protein is selected from the group consisting

of clusterin, T-cell surface glycoprotein CD5 precursor, HSP 84, Ca²⁺ dependent phospholipase A2 precursor, and the ribosomal proteins, S12, S11, L12, L11, S29, S19, L21, L19, L13, L44, and L36.

37. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of clusterin, T-cell surface glycoprotein CD5 precursor, HSP 84, and Ca²⁺ dependent phospholipase A2 precursor and the mRNA is for a protein selected from said group.

38. A method of Claim 1 wherein the protein is selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, CD40, C-MYC, PKC- α , GSTA1, GATA-2, PLGF, ezrin, HGF activator, hepatocyte growth factor-like protein, NCAD, MNDA, LHX1, TIE-1, NCAML1, CD104, CD44, SRC1, NMDA, TKT, ephrin (type A), Sp1, RAB proteins, PKC, TIR, Jak3, EGR-1, TNK1, CAMK IV, HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd153, XPG, XPD, ubiquitin-conjugating enzyme, RAD23, cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, α 3 β 1 integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1, a RAB protein, PKC, TfR, bcl-x and caspase-9.

39. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, CD40, C-MYC, PKC- α , GSTA1, GATA-2, PLGF, ezrin, HGF activator, hepatocyte growth factor-like protein, NCAD, MNDA, LHX1, TIE-1, NCAML1, CD104, CD44, SRC1, NMDA, TKT, ephrin (type A), Sp1, RAB proteins, PKC, TIR, Jak3, EGR-1, TNK1, CAMK IV, HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd153, XPG, XPD, ubiquitin-conjugating enzyme, RAD23, cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, α 3 β 1 integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1, a RAB protein, PKC, TfR, bcl-x and caspase-9. and the mRNA is for a protein selected from said group.

40. A method of Claim 1 wherein the protein is selected from the group consisting of an IL-2 receptor, a gamma c chain of an IL-2 receptor, interferon - γ , and a C-C chemokine.

41. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of an IL-2 receptor, a gamma c chain of an IL-2 receptor, interferon - γ , and a C-C chemokine and the mRNA is for a protein selected from said group.

42. A method of Claim 1 wherein the protein is selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, PKC- α , GSTA1, GATA-2, and PLGF.
- 5 43. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, PKC- α , GSTA1, GATA-2, and PLGF and the mRNA is for a protein selected from said group.
44. A method of Claim 1 wherein the protein is selected from the group consisting of a RAB protein, PKC, and TfR.
- 10 45. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of a RAB protein, PKC, and TfR and the mRNA is for a protein selected from said group.
46. A method of Claim 1 wherein the protein is selected from the group consisting of Jak 3, EGR-1, TNK1, and CAMK IV.
- 15 47. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of Jak 3, EGR-1, TNK1, and CAMK IV and the mRNA is for a protein selected from said group.
48. A method of Claim 1 wherein the protein is selected from the group consisting of HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd 153, XPG, XPD, ubiquitin, conjugating enzyme, RAD 23, and ataxia telangiectasia.
- 20 49. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd 153, XPG, XPD, ubiquitin, conjugating enzyme, RAD 23, and ataxia telangiectasia and the mRNA is for a protein selected from said group.
- 25 50. A method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell a DNA molecule coding for an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule (RNAi), said anti-sense molecule, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for said protein, wherein absent said anti-sense molecule, ribozyme or RNAi nucleic acid, the levels of said protein or its mRNA is less than in a non-Peyer's patch cell.
- 30 51. A method of increasing the extent to which the function of a protein is carried out in a Peyer's patch cell, said method comprising delivering to said cell a nucleic

acid coding for said protein, wherein absent said delivery, the level of said protein or its mRNA is greater in said cell than in a non-Peyer's patch cell.

52. A chimeric protein that comprises two or more segments, each of said segments enhancing a different step in the peptide transport process, said steps selected from the group consisting of binding to a cell, transporting the peptide into the cell, transporting the peptide through the cell, and transporting the peptide out of the cell.

53. A chimeric protein of Claim 52 wherein one of the segments binds to the cell.

54. A chimeric protein of Claim 52 wherein one of the segments is a protein that is more prevalent in a Peyer's patch cell than in a non-Peyer's patch cell.

55. A chimeric protein of Claim 52 wherein the cell is a Peyer's patch cell.

56. A chimeric protein of Claim 55 wherein the cell is an M cell.

57. A method of targeting a composition or delivery vehicle to a Peyer's patch cell said method comprising utilizing a composition or vehicle that contains a protein ligand that will specifically bind to a protein that is up-regulated in Peyer's patch cells.

58. The method of Claim 57 wherein the composition or delivery vehicle comprises a drug or antigen.

59. A method of selecting for a ligand that will selectively bind to a target in a Peyer's patch cell, said method comprising contacting a phage library with a protein that is upregulated in Peyer's patch cells.

60. The method of Claim 59 wherein the protein is attached to a solid substrate.

61. A method of Claim 1 wherein the protein is selected from the group consisting of HGF activator, ezrin, NCAD, MNDA, and LHX1.

62. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of HGF activator, ezrin, NCAD, MNDA, and LHX1, and the mRNA is for a protein selected from said group.

63. A method of Claim 1 wherein the protein is selected from the group consisting of cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, $\alpha 3 \beta 1$ integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1.

64. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, $\alpha 3 \beta 1$ integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1, and the mRNA is for a protein

selected from said group.

- 5 65. A method of promoting enterocyte-M cell conversion, said method comprising orally administering an antigen, antigenic composition, or antigen-carrying particle to a person and either simultaneously with, or prior to, said administration, also orally administering a bacteria, or pro-biotic yogurts, or bacterial component to said person.

AMINO ACID SEQUENCES AND NUCLEOTIDE SEQUENCES CORRESPONDING TO SELECTED
GENBANK ID NUMBERS

5

GENBANK ID: M81750
VERSION M81750.1 GI:895928

10

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15 VMEIKEASSVSDFNQNFVFNRIEIANKT PKISQLYQASGTMVYGLFMLQKKSVMH
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20

GENBANK ID: X59798
VERSION X59798.1 GI:35631

25

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30

GENBANK ID: L27211.1
VERSION L27211.1 GI:558656

35

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45

GENBANK ID: U22398
VERSION U22398.1 GI:790247

40

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55

GENBANK ID: X51521
VERSION X51521.1 GI:31282

60

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65

GENBANK ID: L04143.1
VERSION L04143.1 GI:180574THIS ENTRY IS NOT CONTIGUOUS GENOMIC DNA. IT CONTAINS NUMEROUS PIECES OF
INTRONS.

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GENBANK ID: M32110

55 VERSION M32110.1 GI:189421

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GenBank ID: J04111
VERSION J04111.1 GI:186624

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GENBANK ID: X59932
VERSION X59932.1 GI:30255

20 MSAIQAAWPSGTECIAKYNFHGTAEQDLPFCKGDLTIVAVTKD
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30 GENBANK ID: L29220
VERSION L29220.1 GI:632969

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GENBANK ID: M91815.1
VERSION M91815.1 GI:180169
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55 GENBANK ID: M26708
VERSION M26708.1 GI:190695

60 MSDAAVDTSSSEITTKDLKEKKEVVEEAENGRDAPANGNANEENG
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GENBANK ID: M81757
VERSION M81757.1 GI:337732

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5 GENBANK ID: V00568
DEFINITION HUMAN MRNA ENCODING THE C-MYC ONCOGENE.
VERSION V00568.1 GI:34815

10 MPLNVSTNRRNYDLDYDSVQPYFYCDEEENFYQQQQOSELQPPA
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15 SESGSPSAGHSKPPHSPVLVKRCHVSTHQHNYAAPSTRKDYPAAKRVKLDVVRVLR
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20 GENBANK ID: L29219.1
VERSION L29219.1 GI:632963

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35 GENBANK ID: U49399.1
VERSION U49399.1 GI:1418220

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40 GENBANK ID: XM_039993.2
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4141 GATTCTGAGG CCCTGCCCAA TGAGACTCTA GGTGCCAGTG GATGCCACAG CCCAGCTTGG
4201 CCCTTTCCCT CCAGATCCTG GGTACTGAAA GCCTTAGGGA AGCTGGCCTG AGAGGGGAG
4261 CGGCCCTAAG GGAGTGTCTA AGAACAAGAG CGACCCATTC AGAGACTGTC CCTGAAACCT
4321 AGTACTGCCC CCCATGAGGA AGGAACAGCA ATGGTGTGAG TATCCAGGCT TTGTACAGAG
4381 TGTCTTTCTG TTAGTTTTT ACTTTTTTTG TTTTGTTTT TTAAGACGA AATAAGACCC
4441 CAGGGGAGAA TGGGTGTTGT ATGGGAGGC AAGTGTGGGG GGTCTTCTC CACACCCACT
4501 TTGTCCATTT GCAAATATAT TTTGAAAAA

GENBANK ID: X14798.1

50 SEQUENCE 1:

55 MKAAVDLKPTLTIIKTEKVDLELFPSPDMECADVPLLTSSKEM
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AALCALGKDCFLELAPDFVGDILWEHLEILQKEDVKPYQVNGVNPAYPESRYTSDYFI
SYGIEHAQCVPPEFSEPSFITESYQTLHPISSEELLSLKYENDYPSVILRDLQTD
LQNDYFAIKQEVVTPDNMCMGRSTRGKLGGQDSFESIESYDSCDRLTQSWSSQSFSNS
LQRVPSYDSFDSYPAALPNHKPKGTFKDYVRDRADLNKDKPVPAAALAGYTGSGP
IQLWQFLELLTDKSCQSFISWTGDGWEFKLSDPDEVARRWGRKKNPKMNYEKLRSR
LRYYYDKNIHKTAGKRYVYRFVCDLQSLGTYTPEELHAMLDVKKPDADE

60 SEQUENCE 2;

65 MKAAVDLKPTLTIIKTEKVDLELFPSPDMECADVPLLTSSKEM
MSQALKATFSGFTKEQQLGIPKDPQWTEHVRDWMWAVNEFSLKGVDFQKFCMNG
AALCALGKDCFLELAPDFVGDILWEHLEILQKEDVKPYQVNGVNPAYPESRYTSDYFI
SYGIEHAQCVPPEFSEPSFITESYQTLHPISSEELLSLKYENDYPSVILRDLQTD
LQNDYFAIKQEVVTPDNMCMGRSTRGSGPIQLWQFLELLTDKSCQSFISWTGDGWEF
KLSDPDEVARRWGRKKNPKMNYEKLRSRLRYYYDKNIHKTAGKRYVYRFVCDLQSL

LGYTPEELHAMLDVKPDADE

GENBANK ID: D49547
VERSION D49547.1 GI:710654

MGKDYYQTLGLARGASDEEIKRAYRQALRYHPDKNKEPGAEEK
FKEIAEAYDVLSDPKREIFDRYGEGLKSGSPSGSGGGANGTSFSYTFHGDPHAMF
AEFFGGRNPFDTFFGQRNGEEMDIDDPFSGFPMGMGGFTNVNFGRSRSAQEPARKKQ
DPPVTHDLRVSLLEIYSGCTKKMKISHKRLNPDGKSIRNEDKILTIEVKKGWKEGTKI
TFPKEGDQTSNNIPADIVFVLKDKPHNIFKRDGSDVIYPARISLREALCGCTVNVPTL
DGR TIPVVFKDVIRFGMRKVPGEGLPLPKTPEKRGDLIEFEVIFPERIPQTSRTVL
EQVLPI

GENBANK ID: M11717
VERSION D49547.1 GI:710654

MGKDYYQTLGLARGASDEEIKRAYRQALRYHPDKNKEPGAEEK
FKEIAEAYDVLSDPKREIFDRYGEGLKSGSPSGSGGGANGTSFSYTFHGDPHAMF
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DPPVTHDLRVSLLEIYSGCTKKMKISHKRLNPDGKSIRNEDKILTIEVKKGWKEGTKI
TFPKEGDQTSNNIPADIVFVLKDKPHNIFKRDGSDVIYPARISLREALCGCTVNVPTL
DGR TIPVVFKDVIRFGMRKVPGEGLPLPKTPEKRGDLIEFEVIFPERIPQTSRTVL
EQVLPI

GENBANK ID: X76648
VERSION X76648.1 GI:531404

MAQEFVNCKIQPGKVVFVFIKPTCPYCRAQEILSQLPIKQGLLE
FVDITATNHTNEIQDYLLQLTGARTVPRVFIGKDCIGGCSDLVSLQQSGELLTRLKQI
GALQ

GENBANK ID: NM_011587.1
VERSION NM_011587.1 GI:6755784

MVWWGSSLLPTLFLASHVGSVDLTLLANLRITDPQRFLLTCV
SGEAGAGRSSDPPLLEKDDRIVRTFPPGQPLYLARNGSHQVTLRGFSKPSDLGVFS
CVGGAGARRTRVLYVHNSPGAHLFPDKVTHTVNKGDTAVLSAHVHKEKQTDVIWKNG
SYFNTLDWQADDDGRFQLQONVQPPSSGIYSATYLEASPLGSAFFRLIVRGCGAGRW
GPGCVKDCPGCLHGGVCHDHDGECVCPPGFTGTRCEQACREGRFQSCQEQCPGTAGC
RGLTFCLPDPYGCSCGSGWRGSCQCEACAPDHFADCRLLCQCQNGGTCDFSGCVCP
SGWHGVHCEKSDRIPOILSMATEVEFNIGTMPRINCAAAGNPPFVRGSMKLRKPDGT
LLSTKVIVEPDRTTAEFEVPSLTLDGSGFWECEVSTSGGQDSRRFKVNVKPPVPLTA
PRLAKQSRQLVVSPLVSFSGDGPISVRLHYRPQDSTIAWSAIVDPSENVTLMLNK
PKTGVNVRVQLSRPGEGEGGWGPSALMTTDCPEPLLPWLESWHVEGPDRLRVSWSL
PSVPLSGDGLLRWDGARGQERRENISFPQARTALLTGLTPGTHYQLDVRLYHCTLL
GPASPAHVHLPSPGPPAPRHLHAQALSDSEIQLMWQHPEAPSGPISKYIVEIQVAGG
SGDPQWMDVDRPEETSIIVRGLNASTRYLFRVRASVQGLGDWSNTVEEATLGNLQSE
DPVRESRAAEEGLDQQLVLAVVGSVSATCLTILAALLALVCIRRSCLHRRRTFTYQSG
SGEETILQFSSGTLTLTRRPKPQPEPLSYVLEWEDITFEDLIGENFGQVIRAMIKK
DGLKMNAAIKMLKEYASENDRDFAGELEVLCKLGHHPNIIINLLGACENRGYLYIAIE
YAPYGNLLDFLRKSRVLETDPAFAREHGTASTLSSRQLLRFASDAANGMQYLSEKQFI
HRDLAARNVLVGENLASKIADFGLSRGEEVYVKKTMGRLPVRWMAIESLNYSVYTKS
DVWSFGVLLWEIVSLGGTPYCGMTCAELYEKLPGQYRMEQPRNCDEVYELMRQCWRD
RPERPPFAQIALQLGRMLEARKAYVNMSLFENFTYAGIDATAEEA

GENBANK ID: NM_002867.1
VERSION NM_002867.1 GI:4506368

MASVTDGKHGVKDASDQNFDMFKLLIIGNSSVGKTSFLLRYAD
DTFTPAFVSTVGIDFKVKTVYRHEKRVKLQIWDTAGQERYRTITTAYYRGAMGFILMY
DITNESFNVAQDQWATQIKTYSWDNAQVILVGNKCDMEERVVPTKEGQLLAEQLGFD
FFEASAKENISVRQAFERLVDAICDKMSDSLDTDPSMLGSSKNTRLSDTPPLLQNCSC

GENBANK ID: P27361
NO VERSION DATA

LNENQKLA VKRILSGDCRPLPYILFGPPGTGKTVTIEAVLQVH
FALPDSRILVLCAPSNSAADLVCLRLHESKVLQPATMVRVGHFTHVFDVDEAGQASEPEC
LIPLGLMSDISGQIVLAGDPMQLGSPVIKSLAMAYGLNVSFLERLMSRPAYQRDENAF
5 GACGAHNPLLVTKLVKNYRSHEALLMPLSRLFYHRELEVCA DPTVVTSLLGWEKLPKK
GFPLIFHGVRGSEAREGKSPSWFNPAEAVQVLRVYCCLLAHSISSQVSASDIGVITPYR
KQVEKIRILLRNVDLMDIKVGSVEEFQGEYLVIISTVRSNEDRFEDDRYFLGFLSN
SKRFNVAITRPKALLIVLGNPHVLVRDPCFGALLEYSITNGVYMGCDLPPALQSLQNC
10 GEGVADPSYPVVPPESTGPEKHQEPS

GENBANK ID: NM_002752.1
VERSION NM_002752.1 GI:4506096

MSDSKCD SQFYSVQVADSTFTVLKRYQQLKPIGSGA QGIVCAAF
15 DTVLGISVAVKKLSRPFQNTAKRAYRELVLKCVNHKNII SLNVFTPOKTLEEFQ
DVYLVMEIMDANLCQVIHMELDHERMSYLLYQMLCGIKHLHSAGIIHRDLKPSNIVVK
SDCTLKILDFGLARTACTNFMFTPYVVTRYRAPEVILGMGYKENVDIWSVGCIMGEL
VKGCVIFQGTDHIDQWNKVIEQLGTSPSAEFMKLQPTVRNYVENRPKYPGIKFEELFP
20 DWIFPSESERDKIKTSQARDLLSKMLVIDPKRISVDEALRHPIYITVWYDPAEAEAPP
PQIYDAQLEEREHAIEEWKELIYKEVMDWEERSKNGVVKDQPSAQMQQ

GENBANK ID: M22382.1
VERSION M22382.1 GI:190126

MLRLPTVFRQMRPVSRVLAPHLTRAYAKDVKFGADARALMLQGV
25 DLLADAVAVTMGPKGRTVII EQSWGSPKVT KDGVTVAKSIDLKDKYKNIGAKLVQDVA
NNTNEEAGDGT TATV LARSIAKEGF EKISGANPVEIRRGV MLAVDAVIAELKKQSK
PVTTPEEIAQVATISANGDKEIGNIISDAMKKVGRKGVITVKDGKTLNDELEIEGMK
30 FDRGYISPYFINTSKGQKCEFDQDAYVLLSEKKISSIQSIVPALEIANAHKRPLVIIAE
DVDGEALSTVLNRLKVGLQVAVKAPGFGDNRNQLKDMAIATGGAVFGEGLTLNL
EDVQPHDLGKVGEVIVTKDDAMLLKKGDKAQIEKRIQEIIEQLDVTTSEYEKEKLE
RLAKLSDGVAVLKVGSTSDVEVNEKKDRVTDALNATRAAVEEGIVLGGGCALLRCIPA
LDSLTPANEDQKIGIEIIKRTLKIPAMTIAKNAGVEGSLIVEKIMQSSEVGYDAMAG
35 DFNVMVEKGIIDPTKVVRTALLDAAGVASLLTTAEVVVTEIPKEEKDPGMGAMGMMGG
GMGGGMF

GENBANK ID: U09564.1
DEFINITION HUMAN SERINE KINASE MRNA, COMPLETE CDS.
40 VERSION U09564.1 GI:507212

MERKVLALQARKKRTKAKDKAQRKSETQHRGSAPHSES DLPEQ
EEEILGSDDDDEQEDPN DYCKGGYHLVKIGDLFNGRYHVIRKLGWGHFSTVWLSWDIQQ
KKFVAMKVVKSAEHYTETALDEIRLLKSVRNSDPNDPNREMVVQLLDDFKISGVNGTH
45 ICMVFEVLGHHLKWKIISNYQGLPLPCVKKIIQQVLQGLDYLHTKCRIIHTDIKPEN
ILLSVNEQYIRRLAAEATEWQRSGAPPPSGSAVSTAPQPKPADKMSKNKKKKLKKKQK
ROAELEKRMQEIEMEKESGPGQKRPNKQEESESPVERPLKENPPNKMTEKLEESS
TIGQDQTLMERDTEGGAAEINCNGVIEVINYTONSNNETLRHKEDLHNANDCDVQNLN
QESSFLSLPNGDSSTSQETDSCTPITSEVSDTMVCQSSTVGGQSFSEQHISQLQESIR
50 AEIPCEDEQE QEHNGPLDNKGKSTAGNFLVNPLEPKNAEKLKVKIADLGNACWVHKHF
TEDIQTRQYRSLEVLIGSGYNTPADIWSTACMAFELATGDYLFEPHSGEEYTRDEDHI
ALITIELLGKVPKRLIVAGYSKEFFT KKGDLKHITKLPWGLFEVLVEKYEWSQEAA
GFTDFLLPMELEIPEKRATAAECLRHFWLNS

GENBANK ID: M11507.1
55 VERSION M11507.1 GI:339515

MMDQARSFAFNLFGGEPLSYTRFSLARQVDGDN SHVEMKLAVDE
EENADNNTKANVT KPKRCSGSICYGTIAVIVFLIGFMIGLYGCKGVEPKTECERLA
60 GTESPVREEPGEDFPAARLYWDDLKRLSEKLDSTDTSTIKLLNENSYPREAGSQ
KDENALYVENQFREFFKL SKVWRDQHFKIQVKDSAQNSVIVIDKNGRLVYLVENGG
YVAYSKAATVTGKLVHANFGTKKDFEDLYTPVNGSIVIVRAGKITFAEKVANAESLNA
IGVLTYMDQTKFPVNAELSF FGHHLGTGDPYTPGFPSFNHTQFPSPRS SGLPNIPV
QTISRAAAEKLFGNMEGDCPSDWKT DSTCRMVTSSEKNVCLTVSNVLKEIKILNIFGV
70 IKGFVEPDHYVVVGAQRDAWGPAAKSGVGTALLLKL AQMFSDMVLKDGFPQPSRSIIF
ASWSAGDFGVGATEWLEGYLSSHLKAFYINLDRKAVLGT SNFKVSASPLLYTLEK
65 TMQNVKHPVTGQFLYQDSN WASKVEKLTLDNAAPFLAYSGIPAVSFCFCEDTDYPYL

GTTMDTYKELIERIPELNKVARAAAEVAGQFVIKLTHTDVELNLDYERYNSQLLSFVRD
LNQYRADIKEMGLSLQWLYSARGDFFRATSRLLTDFGNAEKTDRFVMKKLNDRVMRVE
YHFLSPYVSPKESPFHRVFWGSGSHTLPALLENLKLKQNNGAFNETLFRNQLALATW
TIQGAANALSGDVWDIDNEF

5

GENBANK ID: U55017.1
VERSION U55017.1 GI:1297296

10

MESYHKPDQKQLQALKDTANRLRISSIQATTAAGSGHPTSCCSA
AEIMAVLFFHTMRYKSQDPRNPHNDRFVLSKGAAPILYAVWAEAGFLAEALLNLK
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GSVWEAMAFASIYKLDNLVAILDINRLGQSDPAPLQHQMDIYQKRCEAFGWHAIIVDG
HSVEELCKAFGQAKHQPTAIIAKTFKGRGITGVEDKESWHGKPLPKNMAEQIIQEIYS
QIQSKKKILATPPQEDAPSVDIANIRMPSLPSYKVGDKIATRKAYQALAKLGHASDR
IIALDGDGTKNSTFSEIFKKEHPDRFIECYIAEQNMVSIAGCATRNRTVFCSTFAAF
FTRAFDQIIRMAAISESNINLCGSHCGVSIQEDGSPQMALEDLAMFRSVPTSTVFYPSD
GVATEKAVELAANTKGICFIRTSRPENAIYNNNEDFQVGQAKVVLKSKDDQVTVIGA
GVTLHEALAAAELKKKEKINIRVLDPFTIKPLDRKILDSARATKGRILTVEDHYEYEG
GIGEAUVSSAVVGEPIVTHLAVNRVPRSGKPAELLKMGFIORDAIAQAVRGLITKA

20

GENBANK ID: X14034.
VERSION X14034.1 GI:35513

25

MSTTVNVDSIAEYKESQIKRALELGTVMTVFSFRKSTPERRTVQ
VIMETROQVAWSKTADKIEGFLDIMEIKEIRPGKNSKDFERAKAVRQKEDCCFTILYGT
QFVLSTLSLAADSKEDAVNWLSGLKILHQEAMNASTPTTIESWLRKQIYSVDQTRRNS
ISLRELKTLPLINFKVSSAKFLKDKFVEIGAHKDELSFEQFHLFYKMLMFEQQKSIL
DEFKKDSSVFILGNTDRPDASAVYLHDFORFLIHEQQEHWAQDLNKNVREMTKFIDDT
MRETAEPFLFVDEFLTLYLFSRENSIWDEKYDAVDMQDMNPLSHYWISSSHNTYLTGD
QLRSESSPEAYIRCLRMGCRCIELDCWDGPDGKPVYHGWTRTTTKIKFDDVVAIKDH
AFVTSSFPVILSIEEHCSEVQQRHMAKAFKEVFGDLLTKPTEASADQLSPSOLREK
IIIKHKKLGPGRGDVDVNMEDKKDEHKQGGELYMWDSIDQKWTRHYCAIADAKLSFSD
IEQTMEEVEPQDIPPELHFGKEKWFHKKVEKRTSAEKLLOEYCMETGGKDGTFVLVRES
ETFPNDYTLFSWRSGRVQHCRIIRSTMEGGTLKYLLTDNLFRFRMYALIQHYRETHLPC
AEFELRLTDPVPNPNPHESKWPYYDSLRSRGEADMLMRIPRDGAFLIRKREGSDSYAI
TFRARGKVKHCRINRDGRHFVLGTSAYFESLVELVSYEYKHSLYRKMRLRYPTPELL
ERYNTERDINSLYDVSRMYVDPSEINPSMPQRTVKALYDYKAKRSDELSCRGALIH
VSKPEGGWKGVDYGTRIQQYFSPNYVEDISTADFELEKQIIEDNPLGSLCRGILDN
TYNVVKAPOGKNQKSFVFILEPKEQGDPPVEFATDRVEELFEWFQSIREITWKIDSKE
NNMKYWEKNQSIATIELSLVYVCKPTSKTKDNLENPDFREIRSFVETKADSIIRQKPV
DLLKYNQKGLTRVYPKQQRVDSSNYDPFRLWLCGSQMVALNFQADKYMOMNHALFSL
NGRTGYVLQPESMRTEKYDPMPPESQRKILMTLTVKVLGARHLPLKGRSIACPFVEVE
ICGAEGYGNKFKFTTVVNDNGLSPIWAPTQEKVTFEYDPNLAFLRVVVEEDMFSDPN
FLAHATYPIKAVKSGFRSVPLKNGYSEDIELASLLVFCEMRPVLESEEBLYSSCRQLR
RRQEELNNQLFLYDTHQNLNRNANRDALVKEFSVNNHSSCTRNRNATRG

45

GENBANK ID: M27691.1
VERSION M27691.1 GI:181038

50

MTMESGAENQSGDAAVTEAENQQMTVQAQPIATLAQVSMPPA
HATSSAPTTLVQLPNGQTVQVHGVIAAQPSPVQSPQVQVQISTIAESEDQESVD
SVTDSQKRREILSRPSYRKILNDLSSDAPGVPRIEEEKSEETSAPIATTVTVPPI
YQTSSGQYIAITQGGAIQLANNGTDGVQGLQTLMTNAAATQPGTTILQYAQTDDGQQ
ILVPSNQVVVQASGDVQTYQIRTAPTSTIAPGVVMASSPALPTQPAEEAARKREVRL
MKNREAARECRRKKKEYVKLENRVAVLENQNKTLIEELKALKDLYCHKSD

55

GENBANK ID: M18391
VERSION M18391.1 GI:339716

60

MERRWPLGLGLVLLLCAPLPPGARAKEVTLMDSKAQGLGWLL
DPPKDGWSEQQQILNGTPLYMYQDCPMQGRDTHWLRSNWIYRGEASRVHVELQFT
VRDCKSFPGGAGPLGCKETFNLLYMESDQDVGIQLRRPLQKVTTVAADQSFTIRDLA
SGSVKLNVERCSLGRLTRRGLYLAFHNPACVALSVRVFYQRCPETLNGLAQFPDTL
PGPAGLVEVAGTCLPHARASPRPSGAPRMHCSPDGEWLVPVGRCHCEPGYEGGSGEA
CVACPSGSGYRMDMTPHCLTQPGQSTAESEGATICESGHYRAPGEGPQVACTGPPS
APRNLSFSASGTQLSLRWEPPADTGGRRQDVRYSVRCSQCQCTAQDGGPCQPCGVGVHF

65

5 SPGARALTTPAVHVNGLEPYANYTFNVEAONGVSGLGSSGHASTSVSISMGHAESLSG
LSLRLVKKERQLELTWAGSRPRSPGANLTYELHVLNQDEERYQMVLEPRVLLTELQP
DTTYIVRVRLMTPLGPGFSPDHEFRTSPVSRGLTGGEIVAVIFGLLLGAALLLGIL
VFRSRAQRQRQRHVTAFFMWIERTSCAEALCGTSRHTRTLHREPWTLPGGWSNFPS
10 RELDPAWLMVDTVIGEGEFGEVYRGTLRLPSQDCKTVAIKTLKDTSPGGQWNFLREA
TIMGQFSHPHILHLEGVVTKRKPIMIITEFMENAALDAFLREREDQLVPGQLVAMLOG
IASGMNYSNHNHYVRDLAARNILVNQNLCKKVSDFGLTRLDDFDGTETQGGKIPI
RWTAPETIAHRIFTASDVWSFGIVMWEVLSFGDKPYGEMSNQEVMSIEDGYRLPPP
15 VDCPAPLYELMKNCWAYDRARRPHFQKLQAHLEQLLANPHSLRTIANFDPRTVTLRLPS
LSGSDGIPYRTVSEWLESIRMKRYILHFHSAGLDTMECVLELTAEDLTQMGITLPGHQ
KRILCSIQFKD

15 GENBANK ID: X54079.1
VERSION X54079.1 GI:32477

MTERRVPFSLLRGPWDPFDRDWPYHSRLFDQAFGLPRLPEEWSQ
WLGSSSWPGYVRPLPPAAIESPAVAAPAYSRAISRQLSSGVSEIRHTADRWRVSLDVN
HFAPDELTVKTKDGVVEITGKHEERQDEHGYSRCFTRKYTLPPGVDPQTQVSSSLSP
20 GTLTVAPMPKLTQSNITIPVTFESRAQLGGPEAAKSDATAK

20 GENBANK ID: XM_012654.3
VERSION XM_012654.3 GI:14773503

25 MFGVTLWEMFSGGEEPWAGVPPYLILQRLDRARLPRPPLCSRA
LYSLALRCWAPHPSDRPSFSLHLEGLLEAGPSEACVRDVTPEGALRMETGDPITVIE
GSSSFHSPDSTIWKQNGRTFKVGSFPASAVTLADAGGLPATRPVHRGHPCPRSTPR
KHRWRQKEGKSLGCAPSTGPPEEHAPGEDERHFQESGVSSVPRSSSHRGXVQAPLKXR
QARAXAPGTSRPASTPTFIL

30 GENBANK ID: XM_054457.2
VERSION XM_054457.2 GI:18590931

35 MASATDSRYGQKESSDQNFDMFKILIGNSSVGKTSFLFRYAD
DSFTPAFVSTVGIDFKVKTIRNDKRIKLQIWDTAGQERYRTITTAYYRGAMGFILMY
DITNEESFNAVQDWSTQIKTYSWDNAQVLLVGNKCDMEDERVSSERGRQLADHLGFE
FFEASAKDNINVKQTFERLVDVICEKMSSESLDTADPAVTGAKQGPQLSDQQVPPHQDC
AC

40 GENBANK ID: XM_038595.3
VERSION XM_038595.3 GI:18590923

45 MAPPSEETPLIPQRSCSLLSTEAGALHVLLPARGPGPPQRLSFS
FGDHLAEDLCVQAAKASGILPVYHSLFALATEDLSCWFPPSHIFSVEDASTQVLLYRI
RFYFPNWFGLEKCHRFGLRKDLASAILDLPVLEHLFAQHRSDLVSGRLPVGLSLKEQG
ECLSLAVLDLARMAREQAQRPGELLKTVSYKACLPPSLRDLIQGLSFVTRRRIRRTVR
RALRRVAACQADRHSIMAKYIMDLERLDPAGAAETFHVGLPGALGGHDGLGLLRVAGD
GGIAWTQGEQEVLPFCDFPEIVDISIKQAPRVGPAGEHRLVTVRTDNQILEAEFFPG
LPEALSFVALVDGYFRLTTDSQHFFCKEVAPPRLLEEVAEQCHGPITLDFAINKLKTG
50 GSRPGSYVLRSPQDFDLSFLLTVCVQNPGLPDYKGLIRRSPTGTFLLVGLSRPHSS
RELLATCWDGGLHVDGVAVTLTSCIPRPKEKSNLIVVQRGHSPPTSSLVQPQSQYQL
SQMTFFHKIPADSLWHENLGHGSFTKIYRGCRHEVVDGEARKTEVLLKVMADAKHNCM
ESFLEAASLMSQVSYRHLVLLHGVCMAGDSTMVQEFVHLGAIDMYLRKRGHLPASWK
LQVVKQLAYALNYLEDKGLPHGNVSARKVLLAREGADGSPFFIKLSDPGVSPAVLSLE
MLTDRIPWVAPECLREAQTLSEADKWGFATVWEVFSGVTMPIALDPAKKLQFYED
55 RQQLPAPKWTELALLIQCMAYEPVQRPSFRAVIRDLNSLISSDYELLSDPTPGALAP
RDGLWNGAQLYACQDPTIFEEHRLKYISQLGKNFGSVELCRYDPLGDNTGALVAVKQ
LQHSQPDQQRDFQREIQILKALHSDFIVKYRGVSYGPGRQSLRLVMEYLPSCGLRDFL
QRHRARLDASRLLYSSQICKMEYLGSRRCVHRDLAARNILVESEAHVKIADFLAK
LLPLDKDYVVREPGQSPIFWYAPESLSDNIFSRQSDVWSFGVVLYELFTYCDKSCSP
60 SAEFLRMGCERDVPALCRLLELLEEGORLPAPPACPAEVHELMKLCWAPSPQDRPSF
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65 GENBANK ID: NM_002755.2
VERSION NM_002755.2 GI:14589898

5 MPKKKPTPIQLNPAPDGSVNGTSSAETNLEALQKKLEELDE
 QQRKRLEAFLTQKQVGEKDDDFEKISELGAGNGGVVFKVSHKPSGLVMARKLIHLE
 IKPAIRNQIIRELQVLHECNTPYIVGFYGAFFYSDGEISICMEHMDGGSLDQVLKKAGR
 IPEQILGKVSIAVIKGLTYLREKHKIMHRDVKPSNVLNVRGEIKLCLDFGVSGQLIDS
 MANSFVGTRSYMSPERLQGTHYSVQSDIWSMGLSLVEMAVGRYPPIPPDAKELELMFG
 CQVEGDAETPPRPRTGRPLSSYGMDSRPPMAIFELLDYIVNEPPPKLPSPGVFSLEF
 QDFVNKCLIKNPAERADLKQLMVHAFIKRSDAEVDFAGWLCSTIGLNQSTPTHAAG
 V

10 GENBANK ID: NM_001744.1
 VERSION NM_001744.1 GI:4502556

15 MLKVTVPSCSASSCSSTASAAPGTASLVPDYWIDGSNRDALSD
 FFEVESELGRGATSIVYRCKQKGTQKPYALKVLKKTVDKKIVRTEIGVLLRSLHPNII
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 VHRDLKPENLLYATPAPDAPLKIADFGLSKIVEHQVLMKTVCCTPGYCAPEILRGCA
 GPEVDMWSVGIIITYILLCGFEPFYDERGQDMFRILNCEYFISPPWDEVSLNAKDL
 VRKLIIVLDPKKRLTFQALQHPWVTGKAANFVHMDTAQKKLOEFNARRKLKAAVKAVV
 ASSRLGSASSSHGSIQESHKASRDPSPIQDGNEDMKAIPGEKIQGDGAQAQAAVKAQA
 20 ELMKVQALEKVKGADINAEAPKMPKAVEDGIKVADLEEEGLAEELKKTVEEAAAP
 REGQSSAVGFEVPPQDVILPEY

25 GENBANK ID: XM_053461.2
 VERSION XM_053461.2 GI:18553657

30 MASRGATRPNGPNTGNKICQFKLVLLGESAVGKSSLVLRVFKGO
 FHEFQESTIGAAFLTQTVCDDTTVKFEIWDTAGQERYHSLAPMYRGAQAAIVVYDI
 TNEESFARAKNVWVKELOQASPNIVIALSGNKADLANKRAVDQEAQSYADDNSLLEM
 ETSAKTSMNVNEIFMAIAKKLPKNEPQNPGANSARGVDLTEPTQPTRNQCCSN

35 GENBANK ID: U33635.1
 VERSION U33635.1 GI:1016701

40 MGAARGSPARPRRLPLLSVLLPLLGTTQTAIVFIKQPSSQDAL
 QGRRALLRCEVEAPGPVHVWLLDGAPVQDTERFAQGSLSFAAVDPLQDSGTFCQV
 ARDDVTGEEARSANASFNKIEAGPVVLKHPASEAETQPTQVLRCHIDGHPRTY
 QWFRDGTPLSDGQSNHTVSSKERNLTLRPAGPEHSGLYSCAHSFQACSSQNFELS
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30 GENBANK ID: B56529

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3301 TAAATAGATC TTTCAGAGCT GGGTGCTGGG TTTGCCATCT TTTTGTTC TTTGAAAAGC
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3421 AA

GENBANK ID: D26155.1
VERSION D26155.1 GI:505086

45
50
55
60
65

MSTPTDPGAMPHGPGSPGPGSPGPILGSPGPGSPGSPGVHSM
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EDARNPKRKRLMEDELPSWIIKDDAEVERLTCEEEEEKIFGRGSRQRDDVSDAL

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GENBANK ID: M62829.1
VERSION M62829.1 GI:182262

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GENBANK ID: U10421.1
VERSION U10421.1 GI:500756

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GENBANK ID: U08015.1
VERSION U08015.1 GI:500631

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HVPQPSGRTLSLQVANSPIECQSQAQELPLVEKQSTDSYPVVGKKMVLSGHNFLOD
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GENBANK ID: U08191.1
VERSION U08191.1 GI:476273

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GENBANK ID: M55654
 VERSION M55654.1 GI:339491

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GENBANK ID: NM_005568.1
 VERSION NM_005568.1 GI:5031866

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 VSDKEAGSNENDQNLGAKRRGPGTTIKAKQLETLKAAFAATPKPTRHIREQLAQETG
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GENBANK ID: X69111.1
 VERSION X69111.1 GI:32294

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GENBANK ID: NP_000507.1
 VERSION NP_000507.1 GI:4504047

1 MGCLGNSKTE DORNEEKAQR EANKKIEKQL QKDKQVYRAT HRLLLLGAGE SGKSTIVKQM
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 121 NPENQFRVDY ILSVMNVPDF DFPPEFYEAH KALWEDEGVR ACYERSNEYQ LIDCAQYFLD
 181 KIDVIKQADY VPSDQDLRLC RVLTSIGIFET KFQVDKVNHF MFDVGGORDE RRRKWIQCND
 241 VTAIFVVAS SSYNMVIRED NQTNRLQAL NLFKSIWNNR WLRTISVILF LNKQDLLAEK
 301 VLAGKSKIED YFPEFARYTT PEDATPEPGE DPRVTRAKYF IRDEFRLIST ASGDGRHYCY
 361 PHFTCAVDTE NIRRVDNCR DIIQRMHLRQ YELL

GENBANK ID: AAA40889.1
 VERSION AAA40889.1 GI:203357

1 MEQYTANSNS STEQIVVQAG QIQQQQQGGV TAVQLQTEAQ VASASGQQVQ TLQVVQGOPL
 61 MVQVSGGQLI TSTGQPIMVQ AVPGGGQGTI MQVPVSGTQG LQQIQLVPPG QIQIQGGQAV
 121 QVQGGQGGTQ QIIIIQQPQTA VTAGQTQTQQ QIAVQGGQVA QTAEQGTIVY QPVNADGTIL
 181 QQGMITIPAA SLAGAQIVQT GANTNTTSSG QGTVTVTLPV AGNVVNSGGM VMMVPGAGSV
 241 PAIQRIPLPG AEMLEEEPLY VNAKOYHRIL KRRQARAKLE AEGKIPKERR KYLHESRHRH
 301 AMARKRGEGG RFFSPKEKDS PHMQDPNQAD EEAMTQIIRV S

GENBANK ID: B53771
 VERSION B53771 GI:2136296

1 MAWALKLPLA DEVIESGLVQ DFDASLSGIG QELGAGAYSM SDVLALPIFK QEESLPPDN
 61 ENKILPFQYV LCAATSPAVK LHDETLYLN QGQSYEIRML DNRKLGELPE INKLVKSIF
 121 RVVFHDDRLO YTEHQLEGW RWNRPGRDIL DIDIPMSVGI IDPRANPTQL NTVEFLWDEA
 181 KRTSVFIQVH CISTEFTMRK HGGEKGVPPR VQIDTFKENE NGEYTEHLHS ASCQIKVFKP
 241 KGADRKQKTD REKMEKRTPH EKEKYQPSYE TTILTECSPW PEITYVNNNSP SPGFNSSHSS
 301 FSLGEGNGSP NHQPEPPPPV TDNLPTTTP QEAQQLWLRN RFSTFTRLET NFSGADLLKL
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421 NGTFFVYHAI YLEELTAVEL TEKIAQLFSI SPCQISQIYK QGPTGIHVLI SDEMIQNFOE
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GENBANK ID: M92299.1
VERSION M92299.1 GI:184292

MSSYFVNSFSCRYPNPGPDYQLLNYGSGSSLSGSYRDPAMHTGS
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GENBANK ID: M68891.1
VERSION M68891.1 GI:182995

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GENBANK ID: XM_028606.2
VERSION XM_028606.2 GI:15304625

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TISSSGSGESGSPVTSSTTISASLVSSQASSSSFTNANSYSTTTTNSMGINMFT
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GENBANK ID: NP_009077.1
DEFINITION HOMO SAPIENS ZINC FINGER PROTEIN 161 (ZNF161), MRNA.
VERSION NM_007146.1 GI:6005967
CDS 42..1592
/CODON_START=1

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181 CAATAACTCA GAAACCTCAG GGTGCACCAG AACATTAA GGATGCCATT GGGATTAAAA
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361 AAACCCCCAC CACGGTGGTT CCCCTTATCT CTACCATCGC TGGGGACAGC AGCCGAACCT
421 CGTTGGTCTC GACCATGCA GGCATCTTGT CAACAGTCAC TACATCTTCC TCGGGCACCA
481 ACCCCAGTAG CAGTGCCAGC ACCACAGCTA TGCCAGTGAC CCAGTCTGTC AAGAAACCCA
541 GTAAGCCTGT CAAGAAGAAC CATGCTTGTG AGATGTGTGG GAAGGCCTTC CGAGATGTGT
601 ACCATCTCAA TCGACACAAG CTCTCCCAT CAGATGAGAA ACCCTTGAG TGTCTTATT
661 GTAATCAGCG CTTCAAGAGG AAGGACCGGA TGAATTACCA TGTGAGGTCT CATGAAGGAG
721 GCATACCAA ACCCTATACT TGCACTGTTT GTGGGAAAGG CTTCTCAAGG CCTGACCACT
781 TAAGCTGTCA TGTAACACAT GTCCATTCAA CAGAAAGACC CTTCAAATGC CAAACGTGCA
841 CTGCTGCTT TGCCACCAA GACAGACTGC GGACACACAT GGTGCGCCAT GAAGGCAAGG
901 TATCATGTAA CATCTGTGGG AAGCTCCTGA GTGCAGCATA CATACCAGC CACTTAAAGA
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5 1021 GCATGAGTGA AGAGACCACT AACCAAAAGC AGCAGCAGCA GCAGCAGCAG CAACAACAAC
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1141 AAGCTGTTAA AGCAAGGAAG AAAGAAGCTG CTAACCTGTG CCAAACCTCC ACGGCTGCTA
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1441 CCCCAGTGAA TATAGCACAC CCTGTCACTA TCACATCTCC AATGAATCTA CCCACACCTA
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20 2161 GTGATACCAG CTCTGATATG CAAAGCATAT GATAATTTAT CATTCTAACT TCAACGTATA
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2281 GAATATAGCA TTGTAGTTGA CTTTTT

25 GENBANK ID: AAA36598.1
DEFINITION HUMAN STEM CELL PROTEIN (SCL) MRNA, COMPLETE CDS.
VERSION M29038.1 GI:337958
CDS 81..725
/CODON_START=1

30 1 AGTCAGAGTC ACTTCTGTGA AATGGTACTT AGGTAGGCGC GTCCGCTCG GTTACAGCGG
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181 TCTTTGGGGA GCCGGATGCC TTCCCTATGT TCACCACCAA CAATCGAGTG AAGAGGAGAC
241 CTTCCCCCTA TGAGATGGAG ATTACTGATG GTCCCCACAC CAAAGTTGTG CCGCGTATCT
35 301 TCACCAACAG CCGGGAGCGA TGGCGGCAGC AGAATGTGAA CGGGGCTTTT GCCAGCTCC
361 GCAAGCTGAT CCCCACACAT CCCCCGACA AGAAGCTCAG CAAGAATGAG ATCCTCCGCC
421 TGCCATGAA GTATATCAAC TTCTTGGCCA AGCTGCTCAA TGACCAAGGAG GAGGAGGGCA
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40 601 GCGGCAGCTC CCTGGATGGG GCAGCCAGCC CGGACAGCTA CACGGAGGAG CCGCGGCCCA
661 AGCACACGGC CCGCAGCCTC CATCTGCCA TGCTGCCTGC CGCCGATGGA GCCGCCCTC
721 GGTGATGGGT CTGGGCCACC AGGATCAGCC AGGAGGGCGT TCTTAGGCTG CTGGGATGGT
781 GGGCTTCAGG GCAGGTGGGG TGAGAATTGG GCGGCTCTGA AGCAAGGCGG TGGACTTGAA
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45 901 TCCTGTACCA GTAGGAGATC AGAAAAATGG AGCAAGTGG TAGGTACTTT TTGTGAAGAC
961 GGCACGGTCT TCCCTCTTCC CTCAGTCCCA AATCCTTCCC AAGTAAGAGG CTGGAGTTGT
1021 CACTGCTTTT GGCCTGGAGT TTGGGATCCC TGCTTTTCTT AAGACCTGGG GTTGTAGCT
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2041 AATAACCACT GCAAGTCTTT TTGTAAGTG AAGAATCCTT TTGTAGAATG AACCACTGCC
35 2101 CCTTCATTGA TTTCTGTGT CAATCCAGAT GGTGGGATGT GGTTTCTTA AGGTGAGGCC
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5 2221 ATTGTACCCT TAAGTCACCC TAGCCCTCTC CCTCTAGGCT CTGCCCTCGA GGTACAGAGGA
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2341 CCCCCCCTTA TAGTTTGGGC TTCAGCCTAG TGGCTTGTC TCACCATGAT GGGGCCCTAA
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10 2521 TTCAGGCTCC CCAGATCATG TTTTGGTGAA AATTAGGGTT GGTTCCCTTC CAACGTTTGG
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GENBANK ID: M27492.1
VERSION M27492.1 GI:186289

MKVLRLRICFIALLISSLEADKCKEREKIIIVSSANEIDVRPC
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KTVGEGSTSDCDIFVFKVLPEVLEKQCGYKLFYIGRDDYVGEDIVEVINENVKKSRL
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GENBANK ID: L34059
VERSION L34059.1 GI:506409

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FTNQVYNCVDEGSKPGTYVMTITANDADDSTANGMVRYRIVTQTPQSFSQNMFTIN
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RPNLNAINITAADADVHPNIGPYVFELPFVFAVRKNWTITRLNGDYAQLSLRILYLE
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ED

GENBANK ID: M77640
VERSION M77640.1 GI:186053

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5 SQRKHSKRHIHKDHVVVPANTTSVILSGLRPYSSYHLEVQAFNGRSGPASEFTFSTP
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10 HLFKERMFRRHOMAVKNTNGTGRVRLPPAGFATEGWFIGFVSAIILLLLVLLILCFIKRS
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GENBANK ID: M59911.1
VERSION M59911.1 GI:186496

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20 KCVYRGNDLELDDSDDWQTYHNEMCNSNTDYLETGMCQLGTSGGFTQNTVYFGAPGAY
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25 GSLSDHIVLLRARPVINIVHKTLPVPRPAVLDPALCTATSCVQVELCFAYNQSAGNPY
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35 KRQKAEKMSQSPETERLTDDY

GENBANK ID: M81695.1
VERSION M81695.1 GI:487829

MTRTRAALLFTALATSLGFNLDEELTAFRVDSAGFGDSVVQY
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40 SQLLACGPTVHHECGRNMYLTGLCFLGPTQLTQRLPVSROECPRQEQDIVFLIDSG
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45 AIEGTETTSSSSFELEMAQEGFSAVFTPDGPVLGAVGSFTWSGGAFLYPPNMSPTFIN
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50 MQFIPAEIPRSAFECREQVVEQTLVQSNICLYIDKRSKNLLGSRDLQSSVTLDLALD
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RCQVPFSFVQEELEDFTLKGNSLFGWVRQILQKKVSVSVAEITFTDSVYSQLPGQEF
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30 QIAPENGQTTPSPSEK

GENBANK ID: X51841.1
VERSION X51841.1 GI:33910

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5 SOLTSDYITIGFGKFVDKVSVPQTDMRPEKLKEPWPNSDPPFSFKNVISLTEDVDEFNRN
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 10 TRTGSFHIRRGEVGIYQVQLRALEHVDGTHVCQLPEDQKGNHILKPSFSDGLKMDAGI
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 30 PRDYSTLTSVSSHDSRLTAGVPDTPTRLVFSALGPTSLRVSWQEPCEPRPLQGSVVEY
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 35 AQQGGPATAFRVDGDSPESTRTPGLSENVYPKFKVQARTTEGFGPEREGITITESQD
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30 GENBANK ID: XP_030326.1
 VERSION XP_030326.1 GI:14763626

1 MDKFWHAAW GLCLVPLSLA QIDLNITCRF AGVFHVEKNG RYSISRTEAA DLCKAFNSTL
 35 61 PTMAQMEKAL SIGFETCRYG FIEGHVVIPR IHPNSICAN NTGVYILTSN TSQYDTYCFN
 121 ASAPPEEDCT SVTDLPNAFD GPITITIVNR DGTRYVQKGE YRTNPEDIYP SNTDDDDVSS
 181 GSSSERSSTS GGYIFYTFST VHPIPEDDSP WITDSTDRIP ATTLMTSAT ATETATKRQE
 241 TWDWFSWLFL PSESKNHLHT TTQMACTSSN TISAGWEPNE ENEDERDRHL SFGSGIDDD
 301 EDFTSSTST TPRAFDHTKQ NQDWTQWNPS HSNPEVLLQ TTRMTDVRN GTTAYEGNWN
 361 PEAPPLIHH EHHEEETPH STSTIQATPS STEETATQK EQWFGRNWE GYRQTPKEDS
 40 421 HSTGTAAAS AHTSHPMQGR TTPSPEDSSW TDFNFIHPH MGRGHQAGRR MDMSSHSIT
 481 LQPTANPNTG LVEDLORTGP LSMTTQQSNS QSFSTSHGL EEDKDHPPTS TLTSSNRNDV
 541 TGGRRDPNHS EGSTTLLEGY TSHYPHTKES RTFIPVTSK TGSFGVTAVT VGDSNSNVNR
 601 SLSGDDTFFH PSGGSHTHG SESDGHSHGS QEGGANTTSG PIRTPQIPEW LILASLLAL
 45 661 ALILAVCIAV NSRRRCGQKK KLVINSNGA VEDRKPSGLN GEASKSQEMV HLVNKESSET
 721 PDQFMTADET RNLQNVDMKI GV

GENBANK ID: NP_000826.1
 VERSION NP_000826.1 GI:4504129

50 1 MGGALGPALL LTSLFGAWAG LGPGQGEQGM TVAVVFSSSG PPQAQFRARL TPQSFLDLPL
 61 EIQLPLTVGN TNPSSLLTQ ICGLLGAHV HGIVFEDNVD TEAVAQILDF ISSQTHVPIL
 121 SISGSAVVL TPKEPGSAFL QLGVSLEQQL QVLFKVL EY DWSAFVITS LHPGHALFLE
 181 GVRVADASH VSWRLLDVVT LELGPGGPRA RTQRLRLQLD APVFVAYCSR EEAELFAEA
 241 AQAGLVGPGH VWLVPNLALG STDAPPATFP VGLISVVTES WRLSLRQKVR DGVAIALGA
 55 301 HSYWRQHGTL PAPAGDCRVH PGPVSPAREA FYRHLNVTW EGRDFFSFG GYLVOPTMVV
 361 IALNRHRLWE MVGRWEHGV YMKYPVWPY SASLQPVVDS RHLTVATLEE RPFVIVESPD
 421 PGTGGCVPT VPCRRQSNHT FSSGDVAPY KLCKKGFCID ILKKLARVVK MIQEYIDTV
 481 GKHGKVRGV WNGMIGEVY KRADMAIGSL TINEERSEIV DFSVPFVETG ISVMVARSNG
 541 TVSPSAFLEP YSPAVVMMF VMCLTVVAIT VMFEYFSPV SYNQNLTRGK KSGGPAFTIG
 30 601 KSVWLLWALV FNNSVPIENP RGTTSKIMVL VWAFFAVIFL ASYTANLAAF MIQEYIDTV
 661 SGLSDKKFOR PQDQYPPFR GTVPNGSTER NIRSNDYRDM THMVKNQRS VEDALTSKLM
 721 GKLDAFIYDA AVLNYMAGKD EGCKLVITIGS GKVFATTGYG IAMQKDSHWK RAIDLALLQF
 781 LGDGETQKLE TVWLSGICQ EKNEVMSSKL DIDNMAGVFY MLLVAMGLAL LVFAWEHLVY
 841 WKLRHSVPNS SQLDFLLAFS RGIYSCFSGV QSLASPPROA SPDLTASSAQ ASVLKMLQAA
 35 901 RDMVTTAGVS SSLDRATRTI ENWGGGRRAP PPSPCPTPS GPSCLPTPD PPPEPSPTGW
 961 GPPDGGRAL VRRAPQPPGR PPTPGPLSD VSRVSRPAPW EARVPVTRGH CGRHLSASER

1021 PLSPARCHYS SFPRADRSGR PFLPLFPEPP ELEDLPLLGP EQLARREALL HAAWARGSRP
1081 RHASLPSSVA EAFARPSSLP AGCTGPACAR PDGHSACRRL AQAQSMCLPI YREACQEGEQ
1141 AGAPAWQHRQ HVCLHAHAHL PFCWGA VCPH LPPCASHGSW LSGAWGPLGH RGRITLGLGTG
1201 YRDSGGLDEI SSVARGTQGF PGPCTWRRIS SLESEV

GENBANK ID: CAA43045.1

DEFINITION HUMAN CDW40 MRNA FOR NERVE GROWTH FACTOR RECEPTOR-RELATED
B-LYMPHOCYTE ACTIVATION MOLECULE.

VERSION X60592.1 GI:29850

CDS 48..881

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1 GCCTCGCTCG GGCGCCAGT GGTCTGCCG CCTGGTCTCA CCTCGCCATG GTTCGTCTGC
61 CTCTGCAGTG CGTCTCTGG GGTGCTTGC TGACCGCTGT CCATCCAGAA CCACCCACTG
121 CATGCAGAGA AAAACAGTAC CTAATAAACA GTCAAGTGTG TTCTTTGTGC CAGCCAGGAC
181 AGAAACTGGT GAGTGACTGC ACAGAGTTCA CTGAAACGGA ATGCCCTTCT TCGCGTGA
241 GCGAATTCTT AGACACCTGG AACAGAGAGA CACACTGCCA CCAGCACAAA TACTGCGACC
301 CCAACCTAGG GCTTCGGGTC CAGCAGAAGG GCACCTCAGA AACAGACACC ATCTGCACCT
361 GTGAAGAAGG CTGGCACTGT ACGAGTGAGG CCTGTGAGAG CTGTGTCTCT CACCGCTCAT
421 GCTCGCCCGG CTTTGGGGTC AAGCAGATTG CTACAGGGGT TTCTGATACC ATCTGCGAGC
481 CCTGCCAGT CGGCTTCTTC TCCAATGTGT CATCTGCTTT CGAAAAATGT CACCCTTGGA
541 CAAGCTGTGA GACCAAGAC CTGGTTGTGC AACAGGCAGG CACAAACAAG ACTGATGTTG
601 TCTGTGGTCC CCAGGATCGG CTGAGAGCCC TGGTGGTGAT CCCCATCATC TTGGGGATCC
661 TGTTTGCCAT CCTCTTGGT TCAAAAAGGT GGCCAAGAAG CCAACCAATA
721 AGGCCCCCA CCCCAGCAG GAACCCAGG AGATCAATT TCCCGACGAT CTTCTGGCT
781 CCAACACTGC TGCTCCAGT CAGGAGACTT TACATGGATG CCAACCGGTC ACCCAGGAGG
841 ATGGCAAAGA GAGTCGCATC TCAGTGCAGG AGAGACATG AGGCTGCACC CACCCAGGAG
901 TGTGGCCACG TGGGCAACA GGCAGTTGGC CAGAGAGCCT GGTGCTGCTG CTGCAGGGGT
961 GCAGGCAGAA GCGGGGAGCT ATGCCAGTC AGTGCCAGCC CCTC

GENBANK ID: AAB59544.1

DEFINITION HUMAN NERVE GROWTH FACTOR RECEPTOR MRNA, COMPLETE CDS.

VERSION M14764.1 GI:189204

CDS 114..1397

/CODON_START=1

1 GCCGCGGCCA GCTCCGGCGG GCAGGGGGGG CGCTGGAGCG CAGCGCAGCG CAGCCCCATC
61 AGTCCGCAAA GCGGACCGAG CTGGAAGTCG AGCGCTGCCG CGGGAGGCGG GCGATGGGGG
121 CAGGTGCCAC CGGCCGCGCC ATGGACGGGC CGCGCCTGCT GCTGTGTGCT CTTCTGGGGG
181 TGTCCTTGG AGGTGCCAAG GAGGCATGCC CCACAGGCCT GTACACACAC AGCGGTGAGT
241 GCTGCAAAAG CTGCAACCTG GGCAGGGGTG TGGCCAGGCC TTGTGGAGCC AACCCAGCCG
301 TGTGTGAGCC CTGCCCTGGC AGCGTGACGT TCTCCGACGT GGTGAGCGCG ACCGAGCCGT
361 GCAAGCCGTG CACCGAGTGC GTGGGGCTCC AGAGCATGTC GCGCCCGTGC GTGGAGGCCG
421 ACAGCGCCGT GTGCCGCTGC GCCTACGGCT ACTACCAGGA TGAGACGACT GGGCGCTGCG
481 AGGCGTGCCG CGTGTGCGAG GCGGGCTCGG GCCTCGTGTT CTCCTGCCAG GACAAGCAGA
541 ACACCGTGTG CGAGGAGTGC CCCGACGGCA CGTATTCCGA CGAGGCCAAC CACGTGGACC
601 CGTGCCCTGCC CTGCACCGTG TGCAGGACA CCGAGCGCCA GCTCCGCGAG TGACACGCT
661 GGGCCGACGC CGAGTGCAGG GAGATCCCTG GCCGTTGGAT TACACGGTCC ACACCCCCAG
721 AGGGCTCGGA CAGCACAGCC CCCAGCACCC AGGAGCCTGA GGCACCTCCA GAACAAGACC
781 TCATAGCCAG CACGGTGGCA GGTGTGGTGA CCACAGTGAT GGGCAGCTCC CAGCCCGTGG
841 TGACCCGAGG CACCACCGAC AACCTCATCC CTGTCTATTG CTCCATCTTG GCTGCTGTGG
901 TTGTGGGCCT TGTGGCCTAC ATAGCCTTCA AGAGGTGGAA CAGTGCAAG CAGAACAAGC
961 AAGGAGCCAA CAGCCGGCCA GTGAACCAGA CGCCCCACC AGAGGGAGAA AACTCCACA
1021 GCGACAGTGG CATCTCCGTG GACAGCCAGA GCCTGCATGA CCAGCAGCCC CACACGAGA
1081 CAGCCTCGGG CCAGGCCCTC AAGGGTGACG GAGGCCTCTA CAGCAGCCTG CCCCAGCCA
1141 AGCGGGAGGA GGTGGAGAAG CTTCTCAACG GCTCTGCGGG GGACACCTGG CGGCACCTGG
1201 CGGGCGAGCT GGGCTACCAG CCCGAGCACA TAGACTCCTT TACCATGAG GCCTGCCCCG
1261 TTCGCGCCCT GCTTGCAAGC TGGGCCACCC AGGACAGCGC CACACTGGAG GCCCTCTGG
1321 TTGCGCTGCG CCGCATCCAG CGAGCCGACC TCGTGGAGAG TCTGTGCACT GAGTCCACTG
1381 CCACATCCCC GGTGTGAGCC CAACCGGGGA GCGCCCGCCC CGCCCCACAT TCCGACAACC
1441 GATGCTCCAG CCAACCCCTG TGGAGCCGCG ACCCCACCC TTTGGGGGGG GCGCGCTGG
1501 CAGAAGTGG CTCCTCTGGG CAGGACCTCA GAGTCCAGGC CCCAAACCA CAGCCCTGTC
1561 AGTGCAGCCC GTGTGGCCCC TTCACTTCTG ACCACACTTC CTGTCCAGAG AGAGAAGTGC
1621 CCCTGCTGCC TCCCCAACCC TGCCCCCTGCC CCGTACCATT CTCAGGCCAC CTGCCCCCTT

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1681 CTCCCACACT GCTAGGTGGG CCAGCCCCTC CCACCACAGC AGGTGTCATA TATGGGGGGC
1741 CAACACCAGG GATGGTACTA GGGGGAAGTG ACAAGGCCCC AGAGACTCAG AGGGAGGAAT
1801 CGAGGAACCA GAGCCATGGA CTCTACACTG TGAACCTGGG GAACAAGGGT GGCATCCAG
1861 TGGCCTCAAC CCTCCCTCAG CCCCTCTTGC CCCCCACCCC AGCCTAAGAT GAAGAGGATC
1921 GGAGGCTTGT CAGAGCTGGG AGGGGTTTTC GAAGCTCAGC CCACCCCCCT CATTTTGGAT
1981 ATAGGTCAGT GAGGCCAGG GAGAGGCCAT GATTGCCCCA AAGCCAGACA GCAACGGGGA
2041 GGCCAAAGTGC AGGCTGGCAC CGCCTTCTCT AAATGAGGGG CCTCAGGTTT GCCTGAGGGC
2101 GAGGGGAGGG TGGCAGGTGA CCTTCTGGGA AATGGCTTGA AGCCAAAGTCA GCTTTGCCTT
2161 CCACGCTGTC TCCAGACCCC CACCCCTTCC CCACTGCCTG CCCACCCGTG GAGATGGGAT
2221 GCTTGCCCTAG GGCCTGGTCC ATGATGGAGT CAGGTTTGGG GTTCGTGGAA AGGGTGCTGC
2281 TTCCCTCTGC CTGTCCCTCT CAGGCATGCC TGTGTGACAT CAGTGGCATG GCTCCAGTCT
2341 GCTGCCCTCC ATCCCACAT GGACCCGGAG CTAACACTGG CCCCTAGAAAT CAGCCTAGGG
2401 GTCAGGGACC AAGGACCCT CACCTTGCAA CACACAGACA CACGCACACA CACACACAGG
2461 AGGAGAAATC TCACTTTTCT CCATGAGTTT TTTCTCTTGG GCTGAGACTG GATACTGCC
2521 GGGCAGCTG CCAGAGAAGC ATCGGAGGGA ATTGAGGTCT GCTCGGCCGT CTTCACTCGC
2581 CCCCGGGTTT GGCGGGCCAA GGACTGCCGA CCGAGGCTGG AGCTGGCGTC TGTCTTCAAG
2641 GGCTTACACG TGGAGGAATG CTCCCCATC CTCCCTTCC CTGCAACAT GGGGTGGGT
2701 GGGCCAGAA GGTGCGATG AAGAAAAGCG GGCCAGTGTG GGAATGCGGC AAGAAGGAAT
2761 TGACTTCGAC TGTGACCTGT GGGGATTCTT CCCAGCTCTA GACAACCTTG CAAAGGACTG
2821 TTTTTCCTG AGCTTGGCCA GAAGGGGCC ATGAGGCCTC AGTGGACTTT CCACCCCTC
2881 CCTGGCCTGT TCTGTTTTCG CTGAAGTTGG AGTGAGTGTG GCTCCCTCT ATTTAGCATG
2941 ACAAGCCCCA GGCAGGCTGT GCGCTGACAA CCACCGCTCC CCAGCCAGG GTTCCCCAG
3001 CCCTGTGGAA GGGACTAGGA GCACTGTAGT AAATGGCAAT TCTTTGACCT CAACCTGTGA
3061 TGAGGGGAGG AAACCTACCT CTGGCCCCC CACCTGGGCA CCTGGGGAGT GGGACAGAGT
3121 CTGGGTGTAT TTATTTTCTT CTGGGGCCCC TTTTACTCC CCTTGAGCTG AGATGGAACC
3181 GTTTTAGCAT GTGTTTGGTT CTGGGGCCCC TTTTACTCC CCTTGAGCTG AGATGGAACC
3241 CTTTGGCCCC CCAGCTGGGG GCCATGAGCT CCAGACCCC AGCAACCCCTC CTATCACCTC
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VERSION NM_002511.1 GI:4505406
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1 GTGCTGTGAG GCTTGCCCGC GGACAGTAAA CTTGCAGGGG CGAGAGGGAG GGACATCGAT
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301 CACCGTGGGC TTGCTGGGCA ACATCATGCT GGTGAAGATC TTCATCACCACACAGGCCAT
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421 CACCTGCGTC CCGGTGGACG CCTCGCGCTA CTTCTTCGAC GAGTGGATGT TTGGCAAGGT
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1141 TTTCAACAGC CAACCTGCT GTGGGAGGAA GTCTATCAA GAGAGAGGAA CCAGCTACCT
1201 ACTCAGCTCT TCAGCGGTGC GTATGACATC TCTGAAAAGC AATGCTAAGA ACATGGTGAC
1261 CAATTCTGTT TTAATAAATG GGCACAGCAT GAAGCAGGAA ATGGCAATGT GATTTTGGCC
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GENBANK ID: Q00941
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5 CATGACACAGTGCAATTGCGAACGACTTGTCAGTGATGGGGAGTGAGGCCTACTTGGTCGT
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10 GENBANK ID: P05106

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DNA TYPE: CDNA

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20 TGGCAGGCACAGTCACAATCAAAGGTGACCTGGACGATCAGGCTGTCTTGAAGCCACG
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DNA TYPE: CDNA

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35 CCGCTGGCTGCAGACGGGCTGACCCTCCCGGGGGCTGCATTGCTCCTGCTGGGAAGGGCG
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45 EST
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60 GENBANK ID: CAA52348.1
VERSION X74295.1 GI:437781
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121 AGGAACAACCT GGGGCAGCCC CCGGCGGGAG GGCCCGGATG CACACCCCAT CCTGGCTGCT
181 GACGGGCATC CCGAGCTGGG CCCCAGTGGG CATCCAGGGC CAGGCACCGC CTAGGTTCCT
241 ATGTCCAGC CTGCGCTGTG GCTGCCCTCC ATCCCTTCCC CAGAGATGGC TCCTTGGGAT
301 GAAGAGGGTA GAGTGGGCTG CTGGTGTAC ATCAAGAATT TGGCAGGATC GGCTTCTCTCA
361 GGGGCACAGA CCTCTCCAC CCACAAGAAC TCCTCCACCC CAACTTCCCC TTAGAGTGTCT
421 GTGAGATGAG AGTGGGTAAA TCAGGGACAG GGCCATGGGG TAGGGTGAGA AGGGCAGGGG
481 TGTCTGTATG CAAAGGTGGG GAGAAGGATC CTAATCCCTT CCTCTCCCAT TCACCCTGTG
541 TAACAGGACC CCAAGGACCT GCCTCCCGG AAGTGCCTTA ACCTAGAGGG TCGGGGAGGA
601 GGTGTGTCTA CTGACTCAAG GCTGCTCCTT CTCTAGTTTC CCCTCTCATC TGACCTTAGT
661 TTGCTGCCAT CAGTCTAGTG GTTTCGTGGT TTCGTCTATT TATTAATAAA TCGGAACCC

GENBANK ID: M57627
VERSION CAA51942.1 GI:580177
1 MHSSAL

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VERSION AAA52578.1 GI:183364

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1 MWLQSLLLLG TVACISAPAP RSPSPSTQFW EHVNAIQEAR RLLNLSRDTA AEMNETVEVI
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121 ESFKENLKDF LLVIPFDCWE PVQE

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GENBANK ID: AAA58482.1
VERSION AAA58482.1 GI:182669

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1 METNFSIPLN ETEEVLPPEA GHTVLWIFSL LVHGVTFVFG VLGNGLVIWV AGFRMTRTVN
61 TICYLNLALA DFSFSAIPLF RMVSVAMREK WPFASFCLCKL VHVMDINLF VSVYLITIIA
121 LDRICICVLHP AWAQNHRTMS LAKRVMTGLW IFTIVLTLPN FIFWTTISTT NGDYTCIFNF
181 AFWGDTAVER LNVFITMAKV FLILHFIIGF TVPMSIITVC YGIIAAKIHR NHMIKSSRPL
241 RVFAAVVASF FICWFFPYELI GILMAVWLKE MLLNGKYKII LVLINPTSSL AFFNSCLNPI
301 LYVFMGRNFQ ERLIRSLPTS LERALTVEPD SAQTSNTHTT SASPPEETEL QAM

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VERSION P17774 GI:121324

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1 MSESLLVCDV AEDLVEKLK RFRRKETNNA AIIMKIDKDK RLVVLDEELE GISPDELKDE
61 LPERQPRFIV YSYKYQDDG RVSYPLCFIF SSPVGCKPEQ QMMYAGSKNK LVQTAELTKV
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VERSION P51858 GI:1708157

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1 MSRSNRQKEY KCGDLVFAKM KGYPHWPARI DEMPEAAVKS TANKYQVFFF GTHETAFLGP
61 KDLFPYEEK EKFGKPNKRK GFSEGLWEIE NNPTVKASGY QSSQKKSCE EPEPEPEAAE
121 GDGDKKGNAE GSSDEEGKLV IDEPAKEKNE KGALKRRAGD LLEDSPKRPK EAENPEGEEK
181 EAATLEVERP LPMEVEKNST PSEPGSGRGP PQEEEEEEDE EEEATKEDAE APGIRDHESI
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VERSION P10147 GI:127078

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1 MQVSTAALAV LLCTMALCNQ FSASLAADTP TACCFSYTSR QIPQFIADY FETSSQCSKP
61 GVIFLTKRSR QVCADPSEEW VQKYVSDLEL SA

65
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VERSION P13500 GI:126842

1 MKVSAALLCL LLIAATFIPQ GLAQPDAINA PVTCCYNFTN RKISVQRLAS YRRITSSKCP
61 KEAVIFKTIV AKEICADPKQ KVVQDSMDHL DKQTQTPKT

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1 MGVLLTQRTL LSLVLALLFP SMASMAAIGS CSKEYRVLLG QLQKQTDLMQ DTSRLDPYI

5 61 RIQGLDVPKL REHCRERPGA FPSEETLRGL GRRGFLQTLN ATLGCVLHRL ADLEQRLPKA
121 QDLERSGLNI EDLEKLQMAR PNILGLRNNI YCMAQLLDNS DTAEPTKAGR GASQPPTPTP
181 ASDAFQKLE GCRFLHGYHR FMHSVGRVFS KWGESPNRSR RHSPHQALRK GVRTRPSRK
241 GKRLMTRGQL PR

10 GENBANK ID: XP_013053.3
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1 MEKERETLQA WKERVGOELD RVVAFWMEHS HDQEHGGFFT CLGREGRVYD DLKYVWLQGR
61 QVWMYCRLYR TFERFRHAQL LDKAGGGEF LLRYARVAPP GKCAFLVLR DGRPVKVQRT
121 IFSECFYMA MNELWRATGE VRYQTEAVEM MDQIVHWVQE DASGLGRPQL QGAPAAEFMA
181 VPMMLNLVE QLGEADEELA GKYAELGDCW ARRILQHVQR DGQAVLENVS EGGKELPGCL
241 GRQONPGHTL EAGWFLLRHC IRKGDPELRA HVIDKFLLLP FHSGWDPDHG GLFYFQDADN
15 301 FCPTQLEWAM KLWWPHSEAM IAFMLGYSDS GDPVLLRLFY QVAEYTFRQF RDEYGEWFG
361 YLSREGKVAL SIKGGPFKGC FHVPRCLAMC EEMLGALLSR PAPAPSPAPT PACRGAE

20 GENBANK ID: B31848
VERSION B31848 GI:87005

1 MTCKMSQLER NIETIINTFH QYSVKLGHPD TLNQGEFKEL VRKDLQNFLK KENKNEKVIE
61 HIMEDLDTNA DKQLSFEEFI MLMARLTWAS HEKMHEGDEG PGHHHKPGLG EGT

25 GENBANK ID: CAA38698.1
VERSION CAA38698.1 GI:35522

1 MPVMRLFPFCF LQLLAGLALP AVPPQOWALS AGNGSSEVEV VPFQEVWGRS YCRALERLVD.
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30 GENBANK ID: AAA35789.1
VERSION AAA35789.1 GI:181971

1 MNFLLSWVHW SLALLLYLHH AKWSQAAPMA EGGGQNHHEV VKFMDVYQRS YCHPIETLVD
61 IFQYYPDEIE YIFKPSCVPL MRCGGCCNDE GLECVPTES NITMQIMRIK PHQGHIGEM
35 121 SFLQHNKCEC RPKKDRARQE NPCGPCSERR KHLFVQDPQT CKCSCKNTDS RCKARQLELN
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40 AAA66062.1
VERSION AAA66062.1 GI:536898

1 MWKRWLALAL ALVAVAWVRA EEELRSKSKI CANVFCGAGR ECAVTEKGEP TLCIEQCKP
61 HKRPVCGSNG KTYLNHCELH RDACTGSKI QVDYDGHCKE KKSVPSPASP VVCYQSNRDE
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45 181 ITTYPDQENN KLLRGLCVDA LIELSDENAD WKLSFQEFK CLNPSFNPPE KKCALEDETY
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301 KRVSTKEI

50 GENBANK ID: AAA59872.1
VERSION AAA59872.1 GI:398038

1 MGWLPLLLLL TOCLGVPGQR SPLNDFQVLR GTELQHLHA VVPGPWQEDV ADAEECAGRC
61 GPLMDCRAFH YNVSSHGQCL LPWTQHSPT RLRRSGRCDL FQKDYVRTC IMNNGVGYRG
121 TMATTVGGLP CQAWSHKFPN DHKYTPTLRN GLEENFCRNP DGDPPGGPWCY TTDPAVRFQS
55 181 CGIKSCREAA CVWCNGEYR GAVDRTESGR ECQRWDLQHP HQHPFEPGKF LDQGLDDNYC
241 RNPDGSERPW CYTDPQIER EFCDLPRCS EAQPRQEATT VSCFRKGEG YRGTAANTTA
301 GVPCQRWDAQ IPHQHRTPE KYACKDLREN FCRNPDGSEA PWCFTLRPGM RAAFCYQIRR
361 CTDDVRPQDC YHGAGEQYRG TVSKTRKGVQ CQRWSAETPH KPQFTFTSEP HAQLEENFCR
421 NPDGDSHGFW CYTMDPRTPF DYCALRRCAD DQPPSILDPF DQVQFEKCGK RVDRLDQRRS
481 KLRVVGHPG NSPWTVSLRN RQGQHFCCGS LVKEQWILTA RQCFSSCHMP LTGYEVWLGT
60 541 LFQNPQHGEF SLQRPVAKM VCGPSGSQV LKLLERSVT L NQVALICLP PEWYVPPGT
601 KCEIAGWGET KGTGNDTVLN VAFNLVISNQ ECNIKHRGRV RESECTEGL LAPVGACEGD
661 YGGPLACFTH NCWVLEGIII PNRCVARSW PAVFTRVSF VDWIKVMRL G

65 GENBANK ID: P01579
VERSION P01579 GI:124479

1 MKYTSYILAF QLCIVLGS LG CYCQDPYVKE AENLKKYFNA GHSDVADNGT LFLGILKNWK
61 EESDRKIMQS QIVSFYFKLF KNFKDDQSIQ KSVETIKEDM NVKFFNSNKK KRDDFEKLTN
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5 GENBANK ID: XP_035842.1
DEFINITION HOMO SAPIENS SMALL INDUCIBLE CYTOKINE A5 (RANTES) (SCYA5), MRNA.

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15 241 GCCAACCAG AGAAGAAATG GGTTCGGGAG TACATCAACT CTTTGGAGAT GAGCTAGGAT
301 GGAGAGTCTT TGAACCTGAA CTTACACAAA TTGCGCTGTT TCTGCTTGCT CTTGTCTTAG
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421 CACACAGCAG CAGTTACAAA AACCTTCCCC AGGCTGGGAG TGGTGGCTCA CGCCTGTAAT
481 CCCAGCACTT TGGGAGGCCA AGGTGGGTGG ATCACTTGAG GTCAGGAGTT CGAGACCAGC
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30 1141 TGAAAAATCG G

GENBANK ID: CAA72079.1
DEFINITION H. SAPIENS MRNA FOR ESTROGEN SULFOTRANSFERASE.

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45 481 ATTTCTTTCT AATGGTGGCT GGTCATCCAA ATCCTGGATC CTTTCCAGAG TTTGTGGAGA
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55 GENBANK ID: AAA63210.1
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5	541	CAATGACATG	ACTCCAGAGC	AAATGGCTAC	AAATGTGAAC	TGTTCCAGCC	CTGAGCGACA
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25	1801	CCTCAAAGTA	AAATTGAGAA	ATCTTAAAGT	TTTTTTCAAG	TAACATAATC	TATCTTTGTA
	1861	TAATTCATAT	TTGGGAATAT	GGCTTTAAT	AATGTTCTTC	CCACAAATAA	TCATGCTTTT
	1921	TTCTATAGGT	TACAGCATT	AACCTATTTT	TAAGTTGTTT	TTGAACCTTA	TTGTTTGTGT
	1981	ATTTAAGTTT	ATGTTATTTA	TAAAAAATAA	ACCTTAATAA	GCTGTATCTG	TTTCATATGC
30	2041	TTTTAATTTT	AAAGGAATAA	CAAACTGTC	TGGCTCAACG	GCAAGTTTCC	CTCCCTTTTC
	2101	TGACTGACAC	TAAGTCTAG	ACACAGCACT	TGGGCCAGCA	AATCCTGGAA	GCAGACAAAA
	2161	ATAAGAGCCT	GAAGCAATGC	TTACAATAGA	TGCTCACAC	AGAACAATAC	AAATATGTAA
	2221	AAACTCTTTC	ACCACATATT	CTTGCCAATT	AATGGATCA	TATAAGTAAA	ATCATTACAA
	2281	ATATAAGTAT	TTACAGGATT	TTAAAGTTAG	AATATATTGG	AATGCATGGG	TAGAAAAATAT
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	2401	TGTTCAAAAG	GTGGCAGCAC	TGAAGTTGT	TTTCCTGTTA	GATGGCAAGA	GCACAATGCC
	2461	CAAAATAGAA	GATGCAGTTA	AGAAATAAGG	GCCCTGAATG	TCATGAAGGC	TTGAGGTCAG
	2521	CCTACAGATA	ACAGGATTAT	TACAAGGATG	AATTTCCACT	TCAAAGTCT	TTTATTGGCA
40	2581	GATCTTGGA	GCACCTTATA	TGTTCAACCA	TGGGAGGTCA	ATATTTATCT	AAATTTAAAG
	2641	TATGCTAAC	CACCTGTGGT	TTAATTTCAA	AATATTGTCT	ATTCAGTCC	CTTTACATAA
	2701	ATAGTATTTG	GTAATACATT	TATAGATGAG	AGTTATATGA	AAAGGCTAGG	TCAACAAAAA
	2761	CAATAGATT	ATTTAATTTT	CCTGTGGTTG	ACCTATACGA	CCAGGATGTA	GAAAACTAGA
	2821	AAGAAGTCCC	CTTCTCAGA	TATACTCTTG	GGAGAGAGCA	TGAATGGTAT	TCTGAACATAT
45	2881	CACCTGATTC	AAGGACTTTG	CTAGCTAGGT	TTTGAGGTCA	GGCTTCAGTA	ACTGTAGTCT
	2941	TGTGAGCATA	TTGAGGGCAG	AGGAGGACTT	AGTTTTTCAT	ATGTGTTTCC	TTAGTGCCTA
	3001	CGAGACTATC	TGTTTATAAT	CAGTTTTCAG	TGTGAATTCA	CTGAATGTTT	ATAGACAAAA
	3061	GAAAATACAC	ACTAAAATA	ATCTTCATTT	TAAAAGGGTA	AAACATGACT	ATACAGAAAT
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	3181	TGTCTATGGA	GTTATACCTC	CATCAAATTA	CATAGCAATG	CTGAATTAGG	CAAAACCAAC
	3241	ATTTAGTGGT	AAATCCATTC	CTGGTAGTAT	AAGTCACCTA	AAAAAGACTT	CTAGAAATAT
50	3301	GTAATTTAAT	TATTTGTTTT	TCTCCTATTT	TTAAATTTAT	TATGCAAAAT	TTAGAAAAATA
	3361	AAATTTGCTC	TAGTTACACA	CCTTTAGAA	TCTAGAAATAT	TAAAACGTGA	AGGGGCCCTCC
	3421	ATCCCTCTTA	CTCATTGTGA	GTCTAGGAAA	TTGAGATTTT	GATACACCTA	AGGTACAGCA
	3481	GCTGGGTAGA	TATACAGCTG	TCACAAGAGT	CTAGATCAGT	TAGCACATGC	TTTCTACTCT
	3541	TCGATTATTA	GTATTATTAG	CTAATGGTCT	TTGGCATGTT	TTTGTTTTTT	ATTTCTGTGT
55	3601	AGATATAGCC	TTTACATTTG	TACACAAATG	TGACTATGTC	TTGGCAATGC	ACTTCATACA
	3661	CAATGACTAA	TCTATACTGT	GATGATTGGA	CTCAAAGGGA	GAAAAGAAAT	TATGTAGTTT
	3721	TCAATTCGA	TTCTTATTC	CCTTTTGT	ATGAATGGAA	AGCTTTGTGC	AAAATATACA
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	3841	TCTAACAATT	AGA				
60	GENBANK ID: AAA62202.1						
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35	CDS 50..988						
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5 1 GGAACCCGTG GTCCTCCGCT TCATGATTTT CTGCCGTCTC TTGGCAAAAA TGGCAATAA
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 10 421 AGAGAAAATT GAAAGAAAAG GAGAGAAGAA GGAGAAAAAA CAGCAATCAA TAGCTGGAAG
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 541 TGCTAGAAAA CACCCTGATG CAGATTCTTT GTATGTGGAA GAAGTAGATG TCGGAGAAAT
 601 AGCCCCAAGG ACAGTTGTCA GTGGCCTGGT GAATCATGTT CCTCTTGAAC AGATGCAAAA
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 15 721 AGCAATGGTC ATGTGTGCTA GTTCACCAGA GAAATTGAA ATCTTGGCTC CTCCAATGG
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 841 GAATCCTAAG AAGAAGATTT GGGAGCAGAT CCAGCCTGAT CTTACACTA ATGATGAGTG
 901 TGTGGCTACA TACAAAGGAG TTCCCTTTGA GGTGAAAGGG AAGGGAGTAT GTAGGGCTCA
 961 AACCATGAGC AACAGTGGAA TCAAATAAAA TGCTTCCACT ACCAAAAGAC ATTAGAGAAA
 20 1021 ACCTTAAAAG TAATAAGAG AAATATATTT GTCACCT

GENBANK ID: P17936

DEFINITION HUMAN ACIDIC FIBROBLAST GROWTH FACTOR MRNA, 5' END, CLONE
 LAMBDA-MJ36.

CDS 358..>478
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30 1 TCCCCAAGGC TAGGAGGCCA ACCTACTAAC AGGTGGGTGG GTATGGTGTG TGGTTTCACT
 61 CAGTTCTTCT CATGGGGTTT CTCTGAGCTC CATTCATACC AGAAAGGGAG CAGGAGAGAG
 121 AGGACAAGTG GATCCAACAG CCTTCGCTCC AGGGGAATCA GGGCATCGCC TCCTTTTCTG
 181 GGAGGACACT CCCTTCTGAT GGTGAATGGG AACTCCCTTC CTCCTGCAGC AGCCTGCCTG
 241 CAGCTGTCCT GGTAGAACAG TGTGGACATT GCAGAAGCTG TCACTGCCCC AGAAAGAAAG
 301 CACCCACAGG CCAAGGCAAA GAGTCTTGAA AGCGCCACAA GCAGCAGCTG CTGAGCCATG
 361 GCTGAAGGGG AAATCACCAC CTTACAGGCC CTGACCGAGA AGTTTAATCT GCCTCCAGGG
 35 421 AATTACAGA AGCCCCAACT CCTTACTGT AGCAACGGGG GCCACTTCTT GAGGATCC

GENBANK ID:

U76376.1

VERSION U76376.1 GI:1923234

40 MCPCLPHRGRGPPAVCACSAHRLGLRSSAAQLTAARLKALGDEL
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GENBANK ID: Y00638

VERSION Y00638.1 GI:34280

45 MYLWLKLLAFGLDTEVFVTGQSPTSPSTGLTTAKMPSVPLS
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 55 RINGPHERYHLEVEAGNTLVRNESHKNCDFRVKDLQYSTDYTFKAYFHNGDYPGEPFIL
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 35 IAAQGPKLKTIGDFWQMI FQRKVKVIVMLTELKHGDQEICAQYWGEQKQTYGDIEVDL
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5 GENBANK ID: AF001383.1
VERSION AF001383.1 GI:2199534

10 MAEMGSKGVTAGKIASNVQKKLTRAQEKVLQKLGKADETKDEQF
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15 PEIRVNHEPEPAGGATPGATLPKSPSQLRKGPVPPPPKHTPSKEVKQEQLSLFEDT
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20 GENBANK ID: XM 038595.3
VERSION XM_038595.3 GI:18590923

25 MAPPSEETFLIPQRSCSLSTEAGALHVLLPARGPGPPQRLSFS
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45 GENBANK ID: M32292.1
VERSION M32292.1 GI:181492

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10 GENBANK ID: X06318
VERSION X06318.1 GI:35488

15 MADPAAGPPPSEGEESTVRFARKGALRQKNVHEVKNHKFTARFF
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GENBANK ID: J04132.1
VERSION J04132.1 GI:623041

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GENBANK ID: U04313.1
VERSION U04313.1 GI:453368

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GENBANK ID: X68968.1
VERSION X68968.1 GI:452315

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GENBANK ID: L03840.1
VERSION L03840.1 GI:182570

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GENBANK ID: AF043342.1
VERSION AF043342.1 GI:2905633

VRSSSRTPSKPVAHVVANPQAEQQLQWLNRRANALLANGVELR
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GENBANK ID: NM_000735.2
VERSION NM_000735.2 GI:10800407

MDYYRKYAAIFLTVLSVFLHVLHSAIPDVQDCPECTLQENPFFSQ
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GENBANK ID: M83533.1
VERSION M83533.1 GI:178541

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AVFYHGQQLLEYTARLDLWRVQAKEEINEMKELREHNENMLRNILPSHVARHFLEKDR
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GENBANK ID: M31767.1
VERSION M31767.1 GI:181615

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GENBANK ID: X14723

VERSION X14723.1 GI:30250

5 MMKTLLLFVGLLLTWESGQVLGDQTVSDNELQEMSNOGSKYVVK
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NETMMALWEECKPCLKQTCMKFYARVCRSGSLVGRQLEEFNLQSSFFYFWMNGDRID
SLLENDRQQTTHMLDVMQDHFSSRASSIIDELFQDRFFTTREPQDTYHYLPFSLPHRRPHF
FFPKSRIVRSLMPFSPYEPNLFHAMFQFFLEMIHEAQQAMDIFHSPAFQHPPTFIR
10 EGGDDRTVCREIRHNSTGCLRMKDQCKREILSVDCSTNNFSAKLRLRELDDESQVA
ERLTKRYNELLKSYQWKMLNTSSLLLEQLNEQFNWVSRLANLTQGEDQYYLRVTTVASH
TSDSDVPSGVTEVVVKLFDSDPITVTVFVEVSRKNPKFMETVAEKALQEYRKKHREE

GENBANK ID: X04391.1
VERSION X04391.1 GI:37186

15 MPMGSLQPLATLYLLGLMLVASCLGRLSWYDPDFQARLTRSNSKC
QGGQLEVYKDGWHMVCQSWSGRSSKQWEDPSQASKVCQRLNCGVPLSLGPFLVYTPQ
SSIICYGQLGFSFNCSSHRNDMCHSLGLTCLPEQKTPPTTRPPPTTTTPEPTAPRLQ
LVAQSGGQHCAGVVEFYSGSLGGTISYEAQDKTQDLENFLCNNLQCGSFLKHLPETEA
20 GRAQDPGEPREHQPLPIQWKIQNSSCTSLHCFRKKPKQSGRVLALLCSGFQPKVQS
RLVGGSSICEGTVEVRQAQWAAALCDSSSARSSLRWEVCREQQCGSVNSYRVLDAGD
PTSRGLFCPHQKLSQCHELWERNYSYCKKVFTCDPNPAGLAAGTVASIIALVLLV
LLVVCGLPLAYKKLVKKFRQKKQROWIGPTGMNQNSFHRNHTATVRSHAENPTASHVD
NEYSQPPRNSRLSAYPALEGVLHRSSMQPDNSSDDYDLHGAQRL

25 GENBANK ID: S78187.1
VERSION S78187.1 GI:243485

30 MEVPQPEPAPGSALSAPAGVCGGAQRPGHLPGLLLGSHGLLGSFV
RAAASSPVTTLTQTMHDLAGLGSRLTHLSLRASESSLSSESSESSDAGLCMDSP
SPMDPHMAEQTFEQAIQAASRIIRNEQFAIRRFQSMFVRLGHSPVLRNITNSQAPDG
RRKSEAGSGAASSGEDKENDGFVKMPWKPTHPSSTHALAEWASRREAFQRPSSAP
DLMCLSPDRKMEVEELSPLALGRFSLTPAEGDTEEDDGFVDILESDLKDDDAVPPGME
SLISAPLVKLTLEKEEEKDLVMYSKCQRLFRSPSPMPCSVIRPILKRLEPQDRDTPVQN
35 KRRRSVTPPEEQEAEFPKARVLRSLKSLCHDEIENLLSDHRELIGDYSAFLQLQTV
GKHQDLFYISPETMVALLTGKFSNIVDKFVIVDCRYPYEYEGGHIKTAVNPLERDAE
SFLKSPAPIACSLDKRVILIHFCEFSSEGRPMCRFIRERDRAVNDYPSLYYPEMYIL
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40 GENBANK ID: Y00096.1
VERSION Y00096.1 GI:30455

45 MREAAFMYSTAVAIFFVLVAALQGSAPRESPLPYHIPLDPEG
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YIKELPKGFSRHHIIKYEPITVKGNEALVHMEVFQCAPEMDSVPFSGPCDSKMKPD
RLNYCRHVLAALWALGAKAFYYPPEAGLAFGGPGSSRYLRLEVHYHNPLVIEGRNDSSG
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50 FASQLHHTLTGRKVVTVLVRDGREWEIVNQDNHYSFHFQEIIMLKVVSVHPGDVLT
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NNEDVCTCPQASVSQFTSVPWNSFNCDVLKALYSFAPISMHCNKSSAVRFQGEWNQ
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55 GENBANK ID: XM_055551.3
VERSION XM_055551.3 GI:18557356

60 MKETQKSTYYITGESKEQVANSFAVERVRKQGFVVYMTPEIDE
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VTISNRLVSSPCCIVTSTYGWTANMEQIMKAQALRDNSTMGYMAKKHLEINPDHPIM
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EVAAEPESDAVDEIPPLEGDEDAARMEEVD

65 GENBANK ID: M84711.1
VERSION M84711.1 GI:182774

- 5 MAVGKNKRLTKGGKKGAKKKVDPFSKKDWYDVKAPAMFNIRNI
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MDLTRDKMCSMVKKWQTMIEAHVDVKTTDGYLLRLFCVGTCKRNNQIRKTSYAQHQQ
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- 10 GENBANK ID: X53505
VERSION X53505.1 GI:36145
- MAEEGIAAGGVMDVNTALQEVLTALIHDLARGIREAAKALDK
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VVGCSVVVKDYGKESQAKDVIBEYFKCKK
-
- 15 GENBANK ID: X06617
VERSION X06617.1 GI:36143
- MADIQTERAYQKOPTIFONKKRVLLGETGKEKLPRYYKNIGLGF
KTPKEAIEGTYYIDKKCPFTGNVSIIRGILSGVVTMKMQRTIVIRRDYLHYIRKYNRF
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-
- 20 GENBANK ID: M55040.1
VERSION M55040.1 GI:177974
- MRPPQCLLHTPSLASPLLLLLLWLLGGGVGAEGREDAELLVTVR
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LDVYDGRFLVQAERTVLVSMNYRVGAFGLALPGSREAPGNVGLLDQRLALQWQENV
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30 DGDFLSDTPEALINAGDFHGLQVLVGVVDEGSYFLVYGAPGFSKDNESLISRAEFLA
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LPKLLSATDTLDEAERQWKAEFHRWSSYVHWNQFDHYSKQDRCSDL
-
- 35 GENBANK ID: NM_000717.2
VERSION NM_000717.2 GI:9951925
- MRMLLALLALSAAEPSASAESHWCEYVQAESSNYPCLVPVKWGG
40 NCQKDRQSPINIVTTAKVDKLGRRFFSGYDKKQTVTVQNNHGSVMMLLENKASISG
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45 GQRTVIKSGAPGRPLFPWALPALLGPMACLLAGFLR
-
- GENBANK ID: S70587.1
VERSION S70587.1 GI:546848
- 50 MTALFLMSMLFGLACGOAMSFCIPTEYTMHIERRECAYLINT
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CKCGKCNTRYSDCIHEAICTNYCTKPQKSYLVGFSV
-
- 55 GENBANK ID: M34057
VERSION M34057.1 GI:339547
- MDTKLMCLLFFFSPLPLVSNHTGRIKVVFTPSICKVTCTKGSC
QNSCEKGNNTTLLISENGHAADTLTATNFRVVICHLPCMNGQCSSRDCKQCPNFTGK
60 LCQIPVHGASVPKLYQHSQQPGKALGTHVIHSTHTLPLTVTSQQGVVKFPPNIVNIH
VKHEPEASVQIHQVSRIDGPTGQKTKEAQPGQSQVSYQGLFVQKTQTIHSTYSHQOVI
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TFSSCVDPDPVISEEKGPCYRLVSSGRQCMYPLSVHLTKQLCCSVGKAGPHCEKCP
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65 EEPEALTFSSREHGARSAPPEVATAPPEKEIPLSDQEKTKLEPGQPQLSPGISAIHLH
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5 VRYTCICYEGYRFSEQQRKCVDI DECTQVQHLC SQRCENTEGSFLCICPAGFMASEE
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YTRTPDHKHCRDIDECCQGNLCVNGQCKNTEGSFRCTCGQGYQLSAAKDQCEDI DECQ
10 HRHLCAHGQCRNTEGSFQCVCDQGYRASGLDHCEDINECLEDKSVQCRGDCINTAGS
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15 LCDNVLAPNVTKECCCTSGAGWGDNCEIFPCPVLGTAEFTEMCPKGKGFVPAGESSS
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PSSCIDGQC VNTEGSYNCFCTHPMVLDA SEKRCIRPAESNEQIEETDVYQDLCEWHL S
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RDALVDFSEQYTPADPYFIQDRFLNSFEELQAECEGILNGCENGRCVRVQEGYTDCD
15 LDGYHLDTAKMTCFDVNECEDLNRM SLCKNAKCINTDGSYKCLCLPGYVPSDKFNYC
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20 GENBANK ID: AF257099.1
VERSION AF257099.1 GI:8037944

MSDAADVTSSEITTEDLKEKEVVEEAENGRDAPAHGNANEENG
EPEADNEVDEEEEGGEEEGDGEEDGDEDEGAESATGKRAAEDDEDDVDQKQKTD
EDD

25 GENBANK ID: L06505.1
VERSION L06505.1 GI:186799

MPPKFDPNEIKVYYLRCTGGEVGATSALAPKIGPLGLSPKKVGD
30 DIAKATGDWKGRLITVKLTIONRQAQIEVVPASALIIKALKEPPDRRKQKNIKHS
NITFDEIVNIARQMRHRSRLARELSGTIKEILGTAQSVGCNVDGRHPHDIIDDINS
GAV
ECPAS

35 GENBANK ID: X79234.1
VERSION X79234.1 GI:495125

MAQDQGEKENPMRELRIKLCNICVGESGGRLTRA AKVLEQLT
GQTPVFSKARYTVRSFGIRREKIAVHCAVRGAKAEILEKGLKVRELELRKNNFSDT
GNFGFGIQEHIDLGIEYDPSIGIYGLDFYVVLGRPGFSIADKKRRTGCIGAKHRISKE
EAMRWFOQKYDGIILPGK

40 GENBANK ID: X59932.1
VERSION X59932.1 GI:30255

MSAIQAAPSGTECIAKYNFHGTAEQDLFFCKGDVLTIVAVTKD
45 PNWYKAKNKVGREGII PANYVQKREGVKAGTKLSLMPWFHGKITREQAERLLYPETG
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DGLCTRLIKPKVMEGTVA AQDEFYRSGWALNMKELKLLQTIGKEFGDVMGLDYRGNK
VAVKCIKNDATAQAFLAEASVMTQLRHSNLVQLLGVI VEEKGLYIVTEYMAKGS LVD
YLRSGRSVLGGDCLLKFSLDVCEAMEYLEGNFVHRDLAARNVLVSEDNVAKVSDFG
50 LTKEASSTQDTGKLPVKWTAPEALREKKFSTKSDVWSFGILLWEIYSFGRVPYPRIP
LDVVPRVEKGYKMDAPDGCPPAVYEVMMKNCWHLDAAMRPSFLQLREQLHIKTHELHL

55 GENBANK ID: AAA98616.1
VERSION AAA98616.1 GI:178428

1 MQGPWVLLLL GLRLQLSLGI IPVEEENPDF WNRQAAEALG AAKKLQPAQT AAKNLIMFLG
61 DGMGVSTVTA ARILKGQKKD KLGPE TFLAM DRFPYVALSK TYSVDKHVPD SGATATAYLC
60 121 GVKGNFQITIG LSAAARFNQC NTRRGNEVIS VVNRKAKGK SVGVVTTTRV QHASPAGTYA
181 HTVNRNWSYD ADVPASARQE GCQDIATQLI SNMDIDVILG GGRKYMFPMPG TPDPEYPDDY
241 SQGGTRLDGK NLVQEWLAKH QGARYVWNRT ELLQASLDPS VTHLMGLFEP GDMKYEIHRD
301 STLDPSLMM TEAALLLSR NPRGFFLVE GGRIDHGHE SRAYRALTET IMFDDAIERA
361 GQLTSEEDTL SLVTADHSV FSGGGYPLRG SSIFGLAPGK ARDRKAYTVL LYGNPGPYVL
65 421 KDGARPDVTE SESGSPEYRQ QSAVPLDGET HAGEDVAVFA RGPOAHLVHG VQEQTFIAHV
481 MAFAACLEPY TACDLAPPAG TTDAAHGPGS VVPALLPLLA GTLLLLGTAT AP

5 GENBANK ID: XM_041507.1
VERSION XM_041507.1 GI:14737457

10 MGCTVSAEDKAAERSKMIDKNLREDGEKAAREVKLLLLGAGES
GKSTIVKQMKIIHEDGYSEECRQYRAVVSNTIQSIMAIVKAMGNLQIDFADPSRAD
DARQLFALSCTAEQGVLPDDLSGVIRRLWADHGVQACFGRSREYQLNDSAAYYLNDL
ERIAQSDYIPTQQDVLRTVKTGTGIVETHFTFKDLHFKMFDVGGQSRERKKWIHCPEG
VTAIIFCVALSAYDLVLAEDEEMNRMHESMKLFDSICNNKWFTDTSIILFLNKKDLFE
EKITHSPLTICFPEYTGANKYDEAASYIQSKFEDLNKRKDTKEIYTHFTCATDTKNVQ
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15 GENBANK ID: NM_001032.2
VERSION NM_001032.2 GI:13904868

MGHQQLYWSHPRKFGQGSRSRCSNRHGLIRKYGLNMCRCQCFR
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20 GENBANK ID: M22430.1
VERSION M22430.1 GI:190888

25 MKTLLLLAVIMIFGLLQAHGNLVNFHRMIKLTGKEAALS YGFY
GCHCGVGGRGSPKDATDRCCVTHDCCYKRLEKRGCGTKFLSYKFSN SGRITCAKQDS
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30 GENBANK ID: X63527
VERSION X63527.1 GI:36127

35 MSMLRLQKRLASSVLRGCKKKVWLD PNETNEIANANSRQQIRKL
IKDGLIIRKPVTVHSRARC RNTLARRKGRHMGIGRKGTANARMPEKVTWMMRRMIL
RRLRLRYRESKKIDRHMYSLYLKVGKNVFNKRILMEHIHKLKADKARKKLADQAE
ARRSKTKEARKREERLQAKKEEIKTLSKEETKK

40 GENBANK ID: AF099644.1
VERSION AF099644.1 GI:4323527

45 MAQFAFESDLHSLQLDAPINAPPARWQKAKEAGPAPSPMR
AANRSHSAGRTPGRTPGKSSSKVQTPSKPGGDRYIPHRSAAQMEVASFLLSKENQPE
NSQTPTKKEHQKAWALNLNGFDVEAKILRLSGKPQNAPEGYQNRKLVYSQKATPGS
SRKTCRYIPSLPDRILDAPETRNDYYLNLVDWSSGNVLAVALDNSVYLWSASSGDILQ
LLQMEQPEYIYSSVAWIKENYLA VGTSSAEVQLWDVQQOKRLRNMTSHSARVGSLSW
NSYILSSGSRSGHIIHHHDVRVAEHHVATLSGHSQEVCGLRWAPDGRHLASGGNDNLVN
WVPSAPGEGGWVPLQFTTQHQGAVKAWCPWQSNVLTATGGGTSDRHIRIWNVCSGAC
LSAVDAHSQVCSILWSPHYKELISGHGFAQNQLVIWKYPTMAKVAELKGHTSRVLSLT
MSPDGATVASAAADETLRLWRCFELDPARRREREKASAAKSSLIHQGIR

50 GENBANK ID: X51466
VERSION X51466.1 GI:31105

55 MVNFTVDQIRAIMDKKANIRNMSVIAHVDHGKSTLTDSLCKAG
IIASARAGETRFTDTRKDEQERCITIKSTAISLFYELSENDLNFQKSKDGAGFLINL
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60 ATSPGKGLPRTFCQLILDPIFKVFDAIMNFKKEETAKLIEKLDIKL DSEDKDKEGKP
LLKAVMRRWLPAGDALLQMITIHLPSPVTAQKYRCCELLYEGPPDDEAMGIKSCDPKG
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ILMMGRYVEPIEDVPCGNIVGLVGVDQFLVKTGTITTFEHAHNMRVMKFSVSPVVRVA
VEAKNPADLPKLVGLKRLAKSDPMVQCIIEESGEHIIAGAGELHLEICLKDLEEDHA
65 CIPIKKSDPVVSYRET VSEESNVLCLSKSPNKHNRLYMKARFFPDGLAEDIDKGEVSA
RQELKQRARYLAEKYEWDAEARKIWCFCGPDGTGPNILTITKG VQYLNEIKDSVVAG
FQWATKEGALCEENMRGRVFDVHDVTLHADAIHRGGGQI IPTARRCLYASVLT AQPRL
MEPIYLVETIQCPQVVGGIYGLNRRKRGHVFEESQVAGT PMFVVKAYLPVNESFGFTA
DLRSNTGGQAFPCVFDHWQILPGDPDNSSRPSQVVAETRKRKGLKEGI PALDNFLD

GENBANK ID: M15661
VERSION M15661.1 GI:337577

5 MVNVPKTRRTFCKKCGKHQPHKVTQYKKGKDSLYAQGRRRYDRK
QSGYGGQTKPIFRKAKTTKKIVLRLECPEPNCRSKRMLAIKRCKHFELGGDKKRKGQ
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10 GENBANK ID: J04823.1
VERSION J04823.1 GI:1311703

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15 GENBANK ID: NM_001760.2
VERSION NM_001760.2 GI:16950657

20 MELLCCCEGTRHAPRAGDPRLLDQORVLQSLRLLEERYVPRASY
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GPSQTSTPTDVTAIHL

25 GENBANK ID: NM_002625.1
DEFINITION HOMO SAPIENS 6-PHOSPHOFRUCTO-2-KINASE/FRUCTOSE-2,6-BIPHOSPHATASE 1
(PFKFB1), mRNA.
VERSION NM_002625.1 GI:4505744
CDS 80..1495

30 1 GAATTCGGGA CAGGTAGTAA GATAGGAAGT GAGGCCAGGT ACCTTGTGGG CAGTGATGTC
61 ATTCGGGTGCG ACTCCTAAGA TGTCTCCAGA GATGGGAGAG CTCACCCAAA CCAGGTTGCA
35 121 GAAGATCTGG ATTCCACACA GCAGCGGCAG CAGCAGGCTG CAACGGAGAA GGGGCTCATC
181 CATACCCAG TTTACCAATT CCCCACAAT GGTGATCATG GTGGGTTTAC CAGCTCGAGG
241 CAAGACCTAT ATCTCCACAA AGCTCACACG ATATCTCAAC TGGATAGGAA CACCAACTAA
301 AGTGTTAAT TTAGGCCAGT ATCGACGAGA GGCAGTGAGC TACAAGAAGT ATGAATTCTT
361 TCTTCCAGAC AACATGGAAG CCCTGCAAT CAGGAAGCAG TGCGCCCTGG CAGCCCTGAA
421 GGATGTTTAC AACTATCTCA GCCATGAGGA AGGTCATGTT GCGGTTTTTG ATGCCACCAA
40 481 CACTACCAGA GAACGACGGT CACTGATCCT GCAGTTTGCA AAAGAACATG GTTACAAGGT
541 GTTTTTTCAT GAGTCCATT GTAATGACCC TGGCATAATT GCAGAAAACA TCAGGCAAGT
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45 841 CTACCTTTGC CGACATGGCG AGAGTGAAGT CAACATCAGA GGCCGCATCG GAGGTGACTC
901 TGGCCTCTCA GTTCGCGGCA AGCAGTATGC CTATGCCCTG GCCAAGCTCA TTCAGTCCCA
961 GGGCATCAGC TCCCTGAAGG TGTGGACCAG TCGCATGAAG AGGACCATCC AGACAGCTGA
1021 GGCCCTGGGT GTCCCTATG AGCAGTGGAA GGCCCTGAAT GAGATTGATG CGGGTGTCTG
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50 1141 CCAAGATAAA TATCGCTACC GCTATCCCAA GGGAGAGTCC TATGAGGATC TGGTTTCAGCG
1201 TCTGGAGCCA GTGATAATGG AGCTAGAACG ACAGGAGAAT GTACTGGTGA TCTGCCACCA
1261 GGCTGTCATG CGGTGCCCTCC TGGCCTATTT CCTGGATAAA AGTTTCAGATG AGCTTCCATA
1321 TCTCAAGTGC CCTCTGCACA CAGTGCTCAA ACTACTCCT GTGGCTTATG GCTGCAAGT
1381 GGAATCCATC TACCTGAATG TGGAGGCCGT GAACACACAC CGGGAGAAGC CTGAGAATGT
55 1441 GGACATCACC CGGGAACCTG AGGAAGCCCT GGATACTGTC CCAGCCCACT ACTGAGCCCT
1501 TTCCAAGAAG TCAAACTGCC TGTGTCTCTA TCGCCTTCCA CCTTTAGGAA ATGCTATCTT
1561 TGCTCTTCTC CTACTCTGCC TTGGCCTCAC TGAGGCACCC CACTTCCAGT GAAGAAGTCC
1621 TCCGCAACTC CCAACAAGC CTCGCTTGCT GGCCGCAACC AAGGAGCTAT CTAGCTCTGG
1681 AGGAAACTTT CTTTCTTAAT TCCTATTCTC TGACGAATAA AGACTTACTG CCTACAAGAG
60 1741 G

65 GENBANK ID: D00760
DEFINITION HUMAN MRNA FOR PROTEASOME SUBUNIT HC3.
VERSION D00760.1 GI:220023
CDS 1..705
/CODON_START=1

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5 61 CAGATTGAAT ATGCTTTTGGC TGCTGTAGCT GGAGGAGCCC CGTCCGTGGG AATTAAAGCT
121 GCAAAATGGTG TGGTATTAGC AACTGAGAAA AAACAGAAAT CCATTCTGTA TGATGAGCGA
181 AGTGTACACA AAGTAGAACC AATTACCAAG CATATAGGTT TGGTGTACAG TGGCATGGGC
241 CCCGATTACA GAGTGCTTGT GCACAGAGCT CGAAACTAG CTCAACAATA CTATCTTGTG
301 TACCAAGAAC CCATTCTCTAC AGCTCAGCTG GTACAGAGAG TAGCTTCTGT GATGCAAGAA
10 361 TATACTCAGT CAGGTGGTGT TCGTCCATT GGAGTTTCTT TACTTATTG TGGTTGGAAT
421 GAGGGACGAC CATATTTTATT TCAGTCAGAT CCATCTGGAG CTTACTTTGC CTGGAAGGCT
481 ACAGCAATGG GAAAGAACTA TGTGAATGGG AAGACTTTCC TTGAGAAAAG ATATAATGAA
541 GATCTGGAAC TTGAAGATGC CATTCTATACA GCCATCTTAA CCCTAAAGGA AAGCTTTGAA
601 GGGCAAATGA CAGAGGATAA CATAGAAGTT GGAATCTGCA ATGAAGCTGG ATTTAGGAGG
661 CTTACTCCAA CTGAAGTTAA GGATTACTTG GCTGCCATAG CATAACAATG AAGTGACTGA
15 721 AAAATCCAGA ATTTTCAGATA ATCTATCTAC TTAAACATGT TTAAAGTATG TTTTGTTTTG
781 CAGACTTTTT GCATACTTAT TTCTACATGG TTAAATCGA CTGTTTTTAA AATGACACTT
841 ATAAATCCTA ATAAACTGTT AAACCC

GENBANK ID: P10644

20 GENBANK ID: XM_043948.2
DEFINITION HOMO SAPIENS ALDOLASE A, FRUCTOSE-BISPHOSPHATE (ALDOA), MRNA.
VERSION XM_043948.2 GI:18585537
25 CDS 243..1349
/CODON_START=1

1 AAAAACCAGG GCTCCAGAGA ATCAGAACAG CCACCATCAC CGCAGGGAGT CAAGGGAGGA
61 GGGAGATTAG AGAAGGAGCC AGGGAGGGTG GCAGGGAGGC CACGTGATCC GAGTCCCCTC
121 ACCCCTTTCC TTCCCACAGG TCCCTGGCCA AAGATTATAT TCTCTTGACA ACCAAGGGCC
181 TCCGTCTGGA TTTCCAAGGA AGAATTTCTT CTGAAGCACC GGAAGTCTGCT ACTACGAGCA
241 CCATGCCCTA CCAATATCCA GCACTGACCC CGGAGCAGAA GAAGGAGCTG TCTGACATCG
301 CTACCCGCAT CGTGGCACTT GGCAAGGGCA TCCTGGCTGC AGATGAGTCC ACTGGGAGCA
361 TTGCCAAGCG GCTGCAGTCC ATTGGCACCG AGAACACCGA GGAGAACCAG CGCTTCTACC
421 GCCAGCTGCT GCTGACAGCT GACGACCGCG TGAACCCCTG CATTTGGGGT GTCATCCTCT
10 481 TCCATGAGAC ACTCTACAGC AAGGCGGATG ATGGGCGTCC CTTCCTCCAA GTTATCAAA
541 CCAAGGGCGG TGTGTGGGC ATCAAGGTAG ACAAGGGCGT GGTCCTCCCTG GCAGGGACAA
601 ATGGCGAGAC TACCACCCAA GGGTTGGATG GGCTGTCTGA GCGCTGTGCC CAGTACAAGA
661 AGGACGGAGC TGACTTCGCC AAGTGGCGTT GTGTGCTGAA GATTGGGGAA CACACCCCT
721 CAGCCCTCGC CATCATGGAA AATGCCAATG TTCTGGCCCG TTATGCCAGT ATCTGCCAGC
781 AGGTGGGCTT GCAGAATGGC ATTGTGCCCA TCGTGGAGCC TGAGATCCTC CCTGATGGG
841 ACCATGACTT GAAGCGCTGC CAGTATGTGA CCGAGAAGGT GCTGGCTGCT GTCTACAAGG
901 CTCTGAGTGA CCACCACATC TACCTGGAAG GCACCTTGCT GAAGCCCAAC ATGGTCACCC
961 CAGGCCATGC TTGCATCAGC AAGTTTCTCT ATGAGGAGAT TGCCATGGCG ACCGTCACAG
1021 CGCTGCGCGC CACAGTGCCC CCGCTGTCA CTGGGATCAC CTTCCTGTCT GGAGGCCAGA
10 1081 GTGAGGAGGA GGCCTCCATC AACCTCAATG CCATTAACAA GTGCCCCCTG CTGAAGCCCT
1141 GGGCCCTGAC CTTCTCTTAC GGCCGAGCCC TGCAGGCCCT TGCCTGAAG GCCTGGGGCG
1201 GGAAGAAGGA GAACCTGAAG GCTGCGCAGG AGGAGTATGT CAAGCGAGCC CTGGCCAACA
1261 GCCTTGCTTG TCAAGGAAAG TACACTCCGA GCGGTCAGGC TGGGGCTGCT GCCAGCGAGT
1321 CCCTTTCCTG CTCTAACCAC GCCTATTAA GCGAGGTGTT CCCAGGCTGC CCCCACACT
1381 CCAGGCCCTG CCCCCTCCCA CTCTTGAAGA GGAGGCCGCC TCCTCGGGGC TCCAGGCTGG
1441 CTTGCCCGCG CTCTTTCTTC CCTCGTGACA GTGGTGTGTG GTGTCGCTG TGAATGCTAA
1501 GTCCATCACC CTTTCCGACA CACTGCCAAA TAAACAGCTA TTTAAGGGGG

55 GENBANK ID: NM_005175.1
DEFINITION HOMO SAPIENS ATP SYNTHASE, H+ TRANSPORTING, MITOCHONDRIAL FO
COMPLEX, SUBUNIT C (SUBUNIT 9), ISOFORM 1 (ATP5G1), MRNA.
VERSION NM_005175.1 GI:4885080
CDS 120..530
60 /CODON_START=1

1 GGGGAAGCTG AGGGCTGAGA CCAAGGGCTA AAGCTGGGAG GTGAGTCTGT CACCTTGAGC
61 CGGGCGAGCG CTGTGGGCCA AGCAGGGGTT GCAGGGCAGT AGGAGTGCAG ACTGAAAAA
121 TGCAGACCGC CGGGGCATTA TTCATTCTC CAGCTCTGAT CCGCTGTTGT ACCAGGGGTC
181 TAATCAGGCC TGTGTCTGCC TCCTTCTTGA ATAGCCAGT GAATTCATCT AACAGCCTT
241 CCTACAGCAA CTTCCTCTC CAGGTGGCCA GACGGGAGT CCAGACAGT GTTGCTCTCC
65 301 GGGACATTGA CACAGCAGCC AAGTTTATTG GTGCTGGGCG AGCCACAGT GGTGTGGCTG

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361 GTTCAGGGGC TGGCATTGGA ACCGTGTTTG GCAGCTTGAT CATTGGCTAT GCCAGGAACC
421 CGTCTCTCAA GCAGCAGCTC TTCTCCTATG CCATTCTTGG CTTTGGCCCTG TCTGAGGCCA
481 TGGGGCTTTT CTGTTTGATG GTCGCCCTCC TCATCCTCTT CGCCATGTGA GGCTCCATGG
541 GGGGTACCG GCCTGTTGCT ACTGCAACTC CACACCATTG TTGGTGCTGG GGTGTGTTAA
601 GCTTTACCAT TAAACACAAC GTTCTCTAA A

GENBANK ID: M20496.1

DNA LINEAR

DEFINITION HUMAN CATHEPSIN L GENE, COMPLETE CDS.

VERSION M20496.1 GI:809235

CDS 134..1135

/CODON_START=1

15
1 ACCTCCACGT GCCCTGTTTT TCTGGAGGCA CATCCTTGGC CTCTTCCACA GTCCTTGGGT
61 AAATGCTTGG GAGAATAATT TAAATATTTT TATTCTACCA TGGTGGCCCT AATTTTTCAG
121 GGGGCAGTAA GATATGAATC CTACACTCAT CCTTGCTGCC TTTTGCCTGG GAATTCCTC
181 AGCTACTCTA ACATTTGATC ACAGTTTAGA GGCACAGTGG ACCAAGTGGG AGCGCATGCA
241 CAACAGATTA TACGGCATGA ATGAAGAAGG ATGGAGGAGA GCAGTGTGGG AGAAGAACAT
301 GAAGATGATT GAACTGCACA ATCAGGAATA CAGGGAAGGG AAACACAGCT TCACAATGGC
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361 CATGAACGCC TTTGGAGACA TGACCACTGA AGAATTCAGG CAGGTGATGA ATGGCTTTCA
421 AAACCGTAAG CCCAGGAAGG GGAAGTGTG CCAGGAACCT CTGTTTTATG AGGCCCCCAG
481 ATCTGTGGAT TGGAGAGAGA AAGGCTACGT GACTCCTGTG AAGAATCAGG GTCAGTGTGG
541 TTCTTGTGG GCTTTTAGTG CTACTGGTGC TCTTGAAGGA CAGATGTTCG GGAAGTCTG
601 GAGGCTTATC TCACTGAGTG AGCAGATCTT GGTAGACTGC TCTGGGCTC AAGGCAATGA
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661 AGGCTGCAAT GGTGGCCTAA TGGATTATGC TTTCCAGTAT GTTCAGGATA ATGGAGCCCT
721 GGACTCTGAG GAATCCTATC CATATGAGGC AACAGAAGAA TCCTGTAAGT ACAATCCCA
781 GTATTCTGTT GCTAATGACA CCGGCTTTGT GGACATCCCT AAGCAGGAGA AGGCCCTGAT
841 GAAGGCAGTT GCAACTGTGG GGCCCATTTT TGTGCTATT GATGCAGGT ATGAGTCCTT
901 CCTGTCTAT AAAGAAGGCA TTTATTTTGA GCCAGACTGT AGCAGTGAAG ACATGGATCA
30
961 TGGTGTGCTG GTGTTGGGT ACGGATTGA AAGCACAGAA TCAGATAACA ATAAATATTG
1021 GCTGTGTAAG AACAGCTGGG GTGAAGAATG GGGCATGGGT GGCTACGTAA AGATGGCCAA
1081 AGACCGGAGA AACCATTTGT GAATTGCCCTC AGCAGCCAGC TACCCCACTG TGTGAGCTGT
1141 GGACGGTGAT GAGGAAGGAC TTGACTGGGG ATGGCGCATG CATGGGAGGA ATTCTTCAGT
1201 CTACCAGCCC CCGCTGTGTC GGATACACAC TCGAATCATT GAAGATCCGA GTGTGATTTG
35
1261 AATCTGTGA TATTTCACA CTGGTAAATG TTACCTCTAT TTTAATTACT GCTATAAATA
1321 GGTTTATATT ATTGATTCAC TTACTGACTT TGCATTTTCG TTTTAAAG GATGTATAAA
1381 TTTTACCTG TTTAAATAAA ATCG

GENBANK ID: XM 031596.3

DEFINITION HOMO SAPIENS ANNEXIN A4 (ANXA4), MRNA.

VERSION XM 031596.3 GI:18553329

CDS 48..770

/CODON_START=1

45
1 GAAGAACTTC TGCTTGGGTG GCTGAACTCT GATCTTGACC TAGAGTCATG GCCATGGCAA
61 CCAAAGGAGG TACTGTCAA GCTGCTTCAG GATTCAATGC CATGGAAGAT GCCCAGACCC
121 TGAGGAAGGC CATGAAAGGG CTCGGCACCG ATGAAGACGC CATTATTAGC GTCCTTGCCT
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181 ACCGCAACAC CGCCAGCGC CAGGAGATCA GGACAGCCTA CAAGAGCACC ATCGGCAGGG
241 ACTTGATAGA CGACCTGAAG TCAGAACTGA GTGGCAACTT CGAGCAGGTG ATTGTGGGA
301 TGATGACGCC CACGGTGTG TATGACGTGC AAGAGCTGCG AAGGGCCATG AAGGGAGCCG
361 GCACTGATGA GGGCTGCCTA ATTGAGATCC TGGCCTCCCG GACCCCTGAG GAGATCCGGC
421 GCATAAGCCA AACCTACCAG CAGCAATATG GACGGAGCCT TGAAGATGAC ATTCGCTCTG
481 ACACATCGTT CATGTTCCAG CGAGTGCTGG TGTCTCTGTC AGCTGGTGGG AGGGATGAAG
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541 GAAATTATCT GGACGATGCT CTCGTGAGAC AGGATGCCCA GGACCTGTAT GAGGCTGGAG
601 AGAAGAAATG GGGGACAGAT GAGGTGAAAT TTCTAACTGT TCTCTGTTC CGGAACCGAA
661 ATCACTGTT GCATGGTTTG ATGAATACAA AAGGATATCA CAGAAGGATA TTGAACAGAG
721 TATTAAATCT GAAACATCTG GTAGCTTTGA AGATGCTCTG CTGGCTATAG TAAAGTGCAT
781 GAGGAACAAA TCTGCATATT TTGCTGAAAA GCTCTATAAA TCGATGAAGG GCTTGGGCAC
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841 CGATGATAAC ACCCTCATCA GAGTGATGGT TTCTCGAGCA GAAATTGACA TGTGGATAT
901 CCGGGCACAC TTCAAGAGAC TCTATGGAAA GTCTCTGTAC GTGTTTCATCA AGGGTGACAC
961 ATCTGGAGAC TACAGGAAAG TACTGCTTGT TCTCTGTGGA GGAGATGATT AAAATAAAAA
1021 TCCCAGAAGG ACAGGAGGAT TCTCAACACT TTGAATTTT TTAACCTCAT TTTTACAC
1081 TGCTATTATC ATTATCTCAG AATGCTTATT TCCAATTAAA ACGCCTACAG CTGCTCTCTA
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1141 GAATATAGAC TGTCTGTATT ATTATTCACC TATAATTAGT CATTATGATG CTTTAAAGCT
1201 GTACTTGCAT TTCAAAGCTT ATAAGATATA AATGGAGATT TTAAAGTAGA AATAAATATG

5
10
1261 TATTCATGT TTTTAAAAGA TTA CTTTGTGTTT CACAGACATT GAATATATTA
1321 AATTATTCCA TATTTTCTTT TCAGTGAAAA ATTTTAAAG TGGAAAGACTG TTCTAAAATC
1381 ACTTTTTCCT CTAATCCAAAT TTTTAGAGTG GCTAGTAGTT TCTTCATTG AAATGTAAAG
1441 CATCCGGTCA GTAAGAATGC CCATCCAGTT TCTATATTT CATAGTCAAA GCCTTGAAAG
1501 CATCTACAAA TCTCTTTTTT TAGGTTTTGT CCATAGCATC AGTTGATCCT TACTAAGTTT
1561 TTCATGGGAG ACTTCCTTCA TCACATCTTA TGTGAAATC ACTTCTGTA GTCAAAGTAT
1621 ACCAAAACCA ATTTATCTGA ACTAAATTCT AAAGTATGGT TATACAAACC ATATACATCT
1681 GGTTACCAAA CATAAATGCT GAACATTCCA TATTATTATA GTTAATGTCT TAATCCAGCT
1741 TGCAAGTGAA TGGAAAAAAA AATAAGCTTC AAAC TAGGTA TTCTGGGAAT GATGTAATGC
1801 TCTGAATTTA GTATGATATA AAGAAAACCT TTTTGTGCTA AAAATACTTT TTAAATCAA
1861 TTTTGTGAT TGTAGTAATT TCTATTTGCA CTGTGCCTTT CAACCTCCAGA AACATTCTGA
1921 AGATGTACTT GGATTTAATT AAAAAGTTCA CTTTGT

15
GENBANK ID: M22865.1
DEFINITION HUMAN CYTOCHROME B5 MRNA, COMPLETE CDS.
VERSION M22865.1 GI:181226

53..457
CODON_START=1
20
25
30
35
1 CAGCCAGCTC GACGGGGCTG TGTGTGCTGG GCCTGGCTCG CGGCGAACCG AGATGGCAGA
61 GCAGTCGGAC GAGGCCGTGA AGTACTACAC CCTAGAGGAG ATTCAGAAGC ACAACCACAG
121 CAAGAGCACC TGGCTGATCC TGCACCACAA GGTGTACGAT TTGACCAAT TTCTGGAAGA
181 GCATCCTGGT GGGGAAGAAG TTTTAAGGGA ACAAGCTGGA GGTGACGCTA CTGAGAACTT
241 TGAGGATGTC GGGCACTCTA CAGATGCCAG GGAAATGTCC AAAACATTCA TCATTGGGGA
301 GCTCCATCCA GATGACAGAC CAAAGTTAAA CAAGCCTCCG GAAACTCTTA TCACTACTAT
361 TGATTCTAGT TCCAGTTGGT GGACCAACTG GGTGATCCCT GCCATCTCTG CAGTGGCCGT
421 CGCCTTGATG TATCGCCTAT ACATGGCAGA GGACTGAACA CCTCCTCAGA AGTCAGCGCA
481 GGCCGAGCCT GCTTTGGACA CGGGAGAAAA GAAGCCATTG CTAACACTTT CAACTGACAG
541 AAACCTTCAC TTGAAAACAA TGATTTTAAAT ATATCTCTTT CTTTTCTTTC CGACATTAGA
601 AACAAAACAA AAAGAAGTGT CCTTCTGCG CTCAAATTTT TCGAGTGTGC CTTTTATTTC
661 ATCTACTTTA TTTTGATGTT TCCTTAATGT GTAATTTACT TATTATAAGC ATGATCTTTT
721 AAAAATATAT TTGCCTTTTA AAG

35
GENBANK ID: M14362.1
DEFINITION HUMAN T-CELL SURFACE ANTIGEN CD2 (T11) MRNA, COMPLETE CDS.
VERSION M14362.1 GI:179133
CDS 10..1065
/CODON_START=1

40
45
50
55
60
65
1 ACCCCTAAGA TGAGCTTTCC ATGTAAATTT GTAGCCAGCT TCCTTCTGAT TTTCAATGTT
61 TCTTCCAAAG GTGCAGTCTC CAAAGAGATT ACGAATGCCT TGGAAACCTG GGGTGCCTTG
121 GGTCAAGACA TCAACTTGGA CATTCCTAGT TTTCAAATGA GTGATGATAT TGACGATATA
181 AAATGGGAAA AAACCTCAGA CAAGAAAAAG ATTGCACAAAT TCAGAAAAAGA GAAAGAGACT
241 TTCAAGGAAA AAGATACATA TAAGCTATTT AAAAATGGAA CTCTGAAAAT TAAGCATCTG
301 AAGACCGATG ATCAGGATAT CTACAAGGTA TCAATATATG ATACAAAAGG AAAAAATGTG
361 TTGGAAAAAA TATTTGATTG GAAGATTCAA GAGAGGGTCT CAAAACCAAA GATCTCCTGG
421 ACTTGATATCA ACACAACCCT GACCTGTGAG GTAATGAATG GAACTGACCC CGAATTAAAC
481 CTGTATCAAG ATGGGAAACT TCTAAAACCT TCTCAGAGGG TCATCACA CAAGTGGACC
541 ACCAGCCTGA GTGCAAAATT CAAGTGCACA GCAGGGAACA AAGTCAGCAA GGAATCCAGT
601 GTCGAGCCTG TCAGCTGTCC AGAGAAAGGT CTGGACATCT ATCTCATCAT TGGCATATGT
661 GGAGGAGGCA GCCTCTTGAT GGTCTTTGTG GCACTGCTCG TTTTCTATAT CACCAAAGG
721 AAAAAACAGA GGAGTCGGAG AAATGATGAG GAGCTGGAGA CAAGAGCCCA CAGAGTAGCT
781 ACTGAAGAAA GGGGCCGGAA GCGCCACCAA ATTCCAGCTT CAACCCCTCA GAATCCAGCA
841 ACTTCCCAAC ATCCTCCTCC ACCACCTGGT CATCGTTCCC AGGCACCTAG TCATCGTCCC
901 CCGCCTCCTG GACACCGTGT TCAGCACCAG CCTCAGAAGA GGCCTCCTG TCCGTGGGC
961 ACACAAGTTC ACCAGCAGAA AGGCCCGCCC CTCCCAGAC CTCGAGTTCA GCCAAAACCT
1021 CCCCATGGGG CAGCAGAAAA CTCATTGTCC CCTTCTCTA ATTAATAAAG ATAGAAACTG
1081 TATTTTTCAT TAAAAAGCAC TGTGGATTTC TGCCCTCCTG ATGTGCATAT CCGTACTTCC
1141 ATGAGGTGTT TTCTGTGTGC AGAACATTGT CACCTCCTGA GGCTGTGGGC CACAGCCACC
1201 TCTGCATCTT CGAACTCAGC CATGTGGTCA ACATCTGGAG TTTTGTGCT CCTCAGAGAG
1261 CTCCATCACA CCAGTAAGGA GAAGCAATAT AAGTGTGATT GCAAGAATGG TAGAGGACCG
1321 AGCACAGAAA TCTTAGAGAT TTCTGTGCCC CTCTCAGGTC ATGTGTAGAT GCGATAAATC
1381 AAGTGATTGG TGTGCCTGGG TCTCACTACA AGCAGCCTAT CTGCTTAAGA GACTCTGGAG
1441 TTTCTTATGT GCCCTGGTGG AACTTTGCCC ACCATCCTGT GAGTAAAGT GAAATAAAG
1501 CTTTGACTAG

GENBANK ID: XM_087746.1

DEFINITION HOMO SAPIENS SIMILAR TO KIDNEY AMINOPEPTIDASE M; LEUCINE
ARYLAMINOPEPTIDASE 1 (LOC153726), MRNA.

VERSION XM_087746.1 GI:18561749

CDS 262..639

/CODON_START=1

1 GAGTTCCATG CCACCTCCCC GCCCTTTACA GACATGCTAT AAGGTCCCCA GCCCAGTCAC
61 TCCGCAGTGC CTCTCTCTTC CTCCCCATGG ACTATACACA GGCCCTGCTT GTCCTGGAGG
121 AAAGTTTGA CGTCATTATA TAGATCAGGA GACTGAAGTA CTGAAAGGTT AAATGACTTG
181 CCAAAGAATG AGATCTTTTT TTCTAACATT TTACATAATA TCCTCAGAGA AGATCAGGCC
241 CTGGTGACTA GAGCTGTGGC CATGAAGGTG GAAAATTTC AACAAGTGA AATACAGGAA
301 CTCTTTGACA TATTTACTTA CAGCAAGGGA GCGTCTATGG CCCGGATGCT TTCTTGTTTC
361 TTGAATGAGC ATTTATTTGT CAGTGCACTC AAGTCATATT TGAAGACATT TTCCTACTCA
421 AACCGTGAGC AAGATGATCT ATGGAGGCAT TTTCAAATGG CCATAGATGA CCAGAGTACA
481 GTTATTTTGC CAGCAACAAT AAAAAACATA ATGGACAGTT GGACACACCA GAGTGGTTTT
541 CCAGTGATCA CTTTAAATGT GTCTACTGGC GTCATGAAAC AGGAGCCATT TTATCTTGAA
601 AACATTAAAA ATCGGACTCT TCTAACCAGC AATAAGTGAC ACATGGATTG TCCCTATTCT
661 TTGGATAAAA AATGGAACATA CACAACCTTT AGTCTGGCTA GA

GENBANK ID: P31749

DEFINITION DICTYOSTELIUM DISCOIDEUM RAC-ALPHA SERINE/THREONINE KINASE HOMOLOG
MRNA, COMPLETE CDS.

VERSION U15210.1 GI:1000068

CDS 1..1335

/CODON_START=1

1 ATGTCAACAG CACCAATTAA ACATGAAGGT TTCCTCACTA AAGAAGGTGG TGGTTTCAAA
61 AGTTGGAAAA AGAGATGGTT CATTCTCAAA GGTGGTGATT TAAGTTATTA TAAAAACAAA
121 GGTGAACCTG TACCATTAGG AGTTATTCAT TTAATACAT CAGGTCATAT TAAAAATTCT
181 GATCGTAAGA AAAGAGTTAA TGGATTGAA GTACAAACAC CATCACGTAC ATATTTCTTA
241 TGTCAGAGA CAGAGGAAGA ACGTGCAAAA TGGATAGAGA TATTAATTAA TGAAGAGAGA
301 TTATTATTGA ATGGTGGTAA ACAACCAAAG AAATCGGAAA AGGTAGGAGT TGCAGATTTT
361 GAATTATTGA ATTTAGTTGG TAAAGGTAGT TTTGGTAAAG TTATTCAAGT TAGAAAGAAA
421 GATACTGGTG AAGTGTATGC AATGAAAGTT TTATCAAAGA AACATATCGT AGAGCATAAC
481 GAAGTCGAAC ATACATTGAG TGAGCGTAAT ATTCTTCAAA AGATCAATCA CCCATTTTGT
541 GTTAATCTCA ACTACAGTTT TCAACACAGAG GATAAGCTTT ACTTTATCTT GGATTATGTT
601 AATGGTGGTG AGTTATTCTA TCATCTTCAA AAGGACAAAA AGTTTACAGA GGATCGTGTC
661 CGTTATTATG GCGCAGAGAT CGTATTGGCA TTGGAACATT TACATTTGTC GGGTGTATC
721 TATAGAGATT TGAAACCAGA GAATTTACTA CTCACCAACG AGGGTCACAT TTGCATGACC
781 GATTTCCGTC TTTGCAAGA GGGTCTATTG ACACCAACCG ACAAACCTGG TACTTTCTGT
841 GGTACTCTG AATATTAGC ACCGGAAGTA CTTCAAGGCA ATGGTTATGG TAAACAAGTG
901 GATTGGTGGA GTTTTGGTTC TCTCCTCTAT GAAATGCTCA CTGGTTTACC ACCATTCTAC
961 AATCAAGACG TCCAAGAGAT GTATCGTAAG ATCATGATGG AGAAATTATC TTTCCACAT
1021 TTCATTTCTC CAGATGCTCG TTCCCTCTTG GAACAACCTT TGGAAAGAGA TCCTGAAAAA
1081 AGACTTGCCG ATCCAAATCT TATTAAAAGA CATCCTTTCT TCCGTTCCAT CGATTGGGAA
1141 CAATTATTCC AAAAGAATAT TCCACCACCA TTCATTCCAA ATGTTAAAGG TTCTGCTGAT
1201 ACCTCTCAAA TTGATCCAGT TTTCACTGAT GAAGCTCCTT CTTTAACTAT GGCTGGTGAA
1261 TGTGCTTTAA ATCCGCAACA ACAAAGAT TTTGAAGGAT TTACATATGT CGCTGAATCT
1321 GAACATTTAA GATAA

GENBANK ID: NM_000102.2

DEFINITION HOMO SAPIENS CYTOCHROME P450, SUBFAMILY XVII (STEROID
17-ALPHA-HYDROXYLASE), ADRENAL HYPERPLASIA (CYP17), MRNA.

VERSION NM_000102.2 GI:13904854

CDS 61..1587

1 GAGTTGCCAC AGCTCTTCTA CTCACCTGCT GTCTATCTTG CTGCGCGCA CCCAGCCACC
61 ATGTGGGAGC TCGTGGCTCT CTGCTGCTT ACCCTAGCTT ATTTGTTTG GCCCAAGAGA
121 AGGTGCCCTG GTGCCAAGTA CCCCAAGAGC CTCTGTCTCC TGCCCTGGT GGGCAGCCTG
181 CCATTCCTCC CCAGACATGG CCATATGCAT AACAACCTCT TCAAGCTGCA GAAAAAATAT
241 GGCCCATCT ATTCTGTTCT TATGGGCACC AAGACTACAG TGATTGTGCG CCACCACCAG
301 CTGGCCAAGG AGGTGCTTAT TAAGAAGGGC AAGGACTTCT TGGGCGGCG TCAAATGGCA
361 ACTCTAGACA TCGCGTCCAA CAACCGTAAG GGTATCGCCT TCGCTGACTC TGGCGCACAC
421 TGGCAGCTGC ATCGAAGGCT GCGGATGGCC ACCTTTGCCC TGTTCAGGA TGGCGATCAG

5 481 AAGCTGGAGA AGATCATTG TCAGGAAATC AGTACATTGT GTGATATGCT GGCCACCCAC
541 AACGGACAGT CCATAGACAT CTCCTTTCCT GTCTTCGTGG CGGTAACCAA TGTCATCTCC
601 TTGATCTGCT TCAATACCTC CTACAAGAAT GGGGACCCTG AGTTGAATGT CATACAGAAT
661 TACAATGAAG GCATCATAGA CAACCTGAGC AAAGACAGCC TGGTGGACCT AGTCCCCTGG
721 TTGAAGATTT TCCCCAACAA AACCTGGAA AAATTAAAGA GCCATGTTAA AATACGAAAT
781 GATCTGCTGA ATAAATACT TGAAAATTAC AAGGAGAAAT TCCGGAGTGA CTCTATCACC
841 AACATGCTGG ACACACTGAT GCAAGCCAAG ATGAACTCAG ATAATGGCAA TGCTGGCCCA
901 GATCAAGATT CAGAGCTGCT TTCAGATAAC CACATTCTCA CCACCATAGG GGACATCTTT
10 961 GGGGCTGGCG TGGAGACCAC CACCTCTGTG GTTAAATGGA CCCTGGCCTT CCTGCTGCAC
1021 AATCCTCAGG TGAAGAAGAA GCTCTACGAG GAGATTGACC AGAATGTGGG TTTCAGCCGC
1081 ACACCAACTA TCAGTGACCG TAACCGTCTC CTCCTGCTGG AGGCCACCAT CCGAGAGGTG
1141 CTTCCGCTCA GGGCCGTGGC CCCTATGCTC ATCCCCACA AGGCCAACGT TGACTCCAGC
1201 ATCGGTGAGT TTGCTGTGGA CAAGGGCACA GAAGTTATCA TCAATCTGTG GGCCTGTCAT
1261 CACAATGAGA AGGAGTGGCA CCAGCCGAT CAGTTCATGC CTGAGCGTTT CTTGAATCCA
15 1321 CGGGGACCC AGCTCATCTC ACGTCACTA AGCTATTTGC CCTTCGGAGC AGGACCTCGC
1381 TCCTGTATAG GTGAGATCCT GGGCCGCCAG GAGCTCTTCC TCATCATGGC CTGGCTGCTG
1441 CAGAGGTTTC ACCTGGAGGT GCCAGATGAT GGGCAGCTGC CCTCCCTGGA AGGCATCCCC
1501 AAGGTGGTCT TTCTGATCGA CTCTTTCAA GTGAAGATCA AGGTGCGCCA GGCCTGGAGG
1561 GAAGCCACAG CTGAGGGTAG CACCTAAAGG CTGTAATCA CAGCCCTGT CCACCTATG
20 1621 TGGCCCCACA ACACAGATT AGAGATACAA CCCCCACCC TTCTCCGCCA TTCTCCCTA
1681 CTCCCAACCC ACTCTGCCTT CTTTTTCAGC TTGTGGCAAT GCCAGTGATG TGCATAAACA
1741 GTTTTTTTTT TTTC

25 GENBANK ID: NM_001662
DEFINITION HOMO SAPIENS ADP-RIBOSYLATION FACTOR 5 (ARF5), MRNA.
VERSION NM_001662.2 GI:6995999
CDS 37..579
/CODON_START=1

30 1 CCGCGTCGGT GCCCGCGGCC CTCCCGGGC CCCGCCATGG GCCTCACCGT GTCCGCGCTC
61 TTTTCGCGGA TCTTCGGGAA GAAGCAGATG CGGATTCTCA TGGTTGGCTT GGATGCGGCT
121 GGCAAGACCA CAATCCTGTA CAACTGAAG TTGGGGGAGA TTGTCACCAC CATCCCAACC
181 ATAGGCTTCA ATGTAGAAC AGTGAATAT AAGAACATCT GTTTCACAGT CTGGGACGTG
241 GGAGGCCAGG ACAAGATTCG GCCTCTGTGG CCGCACTACT TCCAGAACAC TCAGGGCCTC
35 301 ATCTTTGTGG TGGACAGTAA TGACCGGGAG CGGGTCCAAG AATCTGCTGA TGAACCTCAG
361 AAGATGCTGC AGGAGGACGA GCTGCGGGAT GCAGTGCTGC TGGTATTTGC CAACAAGCAG
421 GACATGCCCA ACGCCATGCC CGTGAGCGAG CTGACTGACA AGCTGGGGCT ACAGCACTTA
481 CGCAGCCGCA CGTGGTATGT CCAGGCCACC TGTGCCACCC AAGGCACAGG TCTGTACGAT
541 GGTCTGGAAT GGCTGTCCCA CGAGCTGTCA AAGCGCTAAC CAGCCAGGGG CAGGCCCTG
40 601 ATGCCCGGAA GCTCCTGCGT GCATCCCCGG GATGACCAGA CTCCCGGACT CCTCAGGCAG
661 TGCCCTTTCC TCCCACTTTT CCTCCCCAT AGCCACAGGC CTCTGCTCCT GCTCCTGCCCT
721 GCATGTTCTC TCTGTTGTTG GAGCCTGGAG CCTGCTCTC TGGGCACAGA GGGGTCCACT
781 CTCTGCTCTG CTGGGACCTA TGGAAAGGGC TTCTTGGCCA AGGCCCTCTC TTCCAGAGGA
841 GGAGCAGGGA TCTGGGTTTC CTTTTTTTTT TCTGTTTGG GTGTACTCTA GGGGCCAGGT
45 901 TGGGAGGGG AAGGTGAGG CTTCGGGTGG TGCTATAATG TGGCACTGGA TCTTGAGTAA
961 TAAATTTGCT GTGGTTTG

50 GENBANK ID: XM_048886.3
DEFINITION HOMO SAPIENS MICROSOMAL GLUTATHIONE S-TRANSFERASE 1 (MGST1), MRNA.
VERSION XM_048886.3 GI:18580621
CDS 89..556
/CODON_START=1

55 1 AGTCCCTGCA TTGCGCGCGA CCCGGCGGCG GGACAGGCTT GCTGCTTCCT CCTCCTCGGC
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121 AGTATTTCAT GCTTTTGCAT CCTATGCAAC AATTATTCTT TCAAAAATGA TGCTTATGAG
181 TACTGCAACT GCATTCTATA GATTGACAAG AAAGGTTTTT GCCAATCCAG AAGACTGTGT
241 AGCATTTGGC AAAGGAGAAA ATGCCAAGAA GTATCTTCGA ACAGATGACA GAGTAGAAGC
301 TGTACGAGA GCCACCTGA ATGACCTTGA AAATATTATT CCATTCTTGT GAATTGGCCT
60 361 CCTGTATTCC TTGAGTGGTC CCGACCCCTC TACAGCCATC CTGCACTTCA GACTATTTGT
421 CGGAGCACGG ATCTACCACA CCATTGCATA TTGACACCC CTTCCTCAGC CAAATAGAGC
481 TTTGAGTTT TTTGTTGGAT ATGGAGTTAC TCTTCCATG GCTTACAGGT TGCTGAAAAG
541 TAAATTGTAC CTGTAAAGAA AATCATACAA CTCAGCATCC AGTTGGCTTT TTAAGAATTC
601 TGTACTTCCA ATTTATAATG AATACTTTCT TAGATTTTAG GTAGGAGGGG AGCAGAGGAA
35 661 TTATGAACCT GGGTAAACCC ATTTGAATA TTAGCATTGC CAATATCCTG TATTCTTGT
721 TTACATTGG ATTAGAAATT TAACATAGTA ATTCTTAAGT CTTTGTCTG ATTTTAAAG

781 TACTTTCTTA TAAATTTGGA TCATGTTATG ATTTGTAACA TTCACACAAC ACCTCACTTT
841 TGAATCTATA AAAGAATTGC ACGTATGAGA AACCTATATT TCAATACTGC TGAACACAGAC
901 ATGAAATAAA GAATTTAAAG AATG

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GENBANK ID: X02162
DEFINITION HUMAN MRNA FOR APOLIPOPROTEIN AI (APO AI)=.
VERSION X02162.1 GI:28771
CDS 87..890
/CODON_START=1

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1 GAATTCAAAA AAAAAAGAAA AAAAAAAAAA AAAAAAAAAA AAAAAAAGAG AGACTGCGAG
61 AAGGAGGTCC CCCACGGCCC TTCAGGATGA AAGCTGCGGT GCTGACCTTG GCCGTGCTCT
121 TCCTGACGGG GAGCCAGGCT CGGCATTTC TGGCAGCAAGA TGAACCCCTT CAGAGCCCTT
181 GGGATCGAGT GAAGGACCTG GCCACTGTGT ACGTGGATGT GCTCAAAGAC AGCGGCAGAG
241 ACTATGTGTC CCAAGTTTGAA GGCTCCGCCT TGGGAAAACA GCTAAACCTA AAGCTCCTTG
301 ACAACTGGGA CAGCGTGACC TCCACCTTCA GCAAGCTGCG CGAACAGCTC GGCCCTGTGA
361 CCCAGGAGTT CTGGGATAAC CTGGAAAAGG AGACAGAGGG CCTGAGGCAG GAGATGAGCA
421 AGGATCTGGA GGAGGTGAAG GCCAAGGTGC AGCCCTACCT GGACGACTTC CAGAAGAAGT
481 GGCAGGAGGA GATGGAGCTC TACCGCCAGA AGGTGGAGCC GCTGCGCGCA GAGCTCCAAG
541 AGGGCGCGCG CCAGAAGCTG CACGAGCTGC AAGAGAAGCT GAGCCCACTG GGCAGGAGGA
601 TGC GCGACCG CGCGCGCGCC CATGTGGACG CGCTGCGCAC GCATCTGGCC CCCACAGCG
661 ACGAGCTGCG CCAGCGCTTG GCCGCGCGCC TTGAGGCTCT CAAGGAGAAC GGC GCGGCCA
721 GACTGGCCGA GTACCAAGCC AAGGCCACCG AGCATCTGAG CACGCTCAGC GAGAAGGCCA
781 AGCCCGCGCT CGAGGACCTC CGCCAAGGCC TGCTGCCCGT GCTGGAGAGC TTCAAGGTCA
841 GCTTCTGAG CGCTCTCGAG GAGTACACTA AGAAGCTCAA CACCCAGTGA GGC GCGGCCG
901 GCCGCCCCC TTCCCGGTGC TCAGAATAAA CGTTTCCAAA GTGGGAAAAA AAAAAAAG
961 AATTC

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GENBANK ID: XM_007441.1
DEFINITION HOMO SAPIENS PRESENILIN 1 (ALZHEIMER DISEASE 3) (PSEN1), MRNA.
VERSION XM_007441.1 GI:11435041
CDS 249..1652
/CODON_START=1

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1 TGGGACAGGC AGCTCCGGGG TCCGCGGTTT CACATCGGAA ACAAAACAGC GGCTGGTCTG
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121 CTGGGAGCCT GCAAGTGACA ACAGCCTTTG CGGTCCCTAG ACAGCTTGGC CTGGAGGAGA
181 ACACATGAAA GAAAGAACCT CAAGAGGCTT TGTCTTCTGT GAAACAGTAT TTCTATACAG
241 TTGCTCCAAT GACAGAGTTA CTTGCACCGT TGTCTTACTT CCAGAATGCA CAGATGTCTG
301 AGGACAACCA CCTGAGCAAT ACTGTACGTA GCCAGAATGA CAATAGAGAA CGGCAGGAGC
361 ACAACGACAG ACGGAGCCTT GGCCACCCCTG AGCCATTATC TAATGGACGA CCCGAGGTA
421 ACTCCCGGCA GGTGTGGAG CAAGATGAGG AAGAAGATGA GGAGCTGACA TTGAAATATG
481 GCGCCAAGCA TGTGATCATG CTCCTTGTCC CTGTGACTCT CTGCATGGTG GTGGTCGTGG
541 CTACCATTA GTCAGTCAGC TTTTATACCC GGAAGGATGG GCAGCTAATC TATACCCCAT
601 TCACAGAAGA TACCGAGACT GTGGGCCAGA GAGCCCTGCA CTCAATTCTG AATGCTGCCA
661 TCATGATCAG TGTCAATTGT GTCATGACTA TCCTCCTGGT GGTCTGTAT AAATACAGGT
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781 CATTCAATTA CTTGGGGGAA GTGTTTAAAA CCTATAACGT TGCTGTGGAC TACATTACTG
841 TTGCACTCCT GATCTGGAAT TTTGGTGTGG TGGGAATGAT TTCCATTAC TGGAAAGGTC
901 CACTTCGACT CCAGCAGGCA TATCTCATTA TGATTAGTGC CCTCATGGCC CTGGTGTGTTA
961 TCAAGTACCT CCCTGAATGG ACTGCGTGGC TCATCTTGGC TGTGATTICA GTATATGATT
1021 TAGTGGCTGT TTTGTGTCG AAAGGTCCAC TTCGTATGCT GGTGAAACA GCTCAGGAGA
1081 GAAATGAAAC GCTTTTCCA GCTCTCATTT ACTCCTCAAC AATGGTGTGG TTGGTGAATA
1141 TGGCAGAAGG AGACCCGGAA GCTCAAAGGA GAGTATCCAA AAATCCAAAG TATAATGCAG
1201 AAAGCACAGA AAGGGAGTCA CAAGACACTG TTGCAGAGAA TGATGATGCG GGGTTCAGTG
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1321 GAGCTGCTGT CCAGGAACCT TCCAGCAGTA TCCTCGCTGG TGAAGACCCA GAGGAAAGGG
1381 GAGTAAACT TGGATTGGGA GATTTTCATTT TCTACAGTGT TCTGGTTGGT AAAGCCTCAG
1441 CAACAGCCAG TGGAGACTGG AACACAACCA TAGCCTGTTT CGTAGCCATA TTAATTGGTT
1501 TGTGCCTTAC ATTATTACTC CTTGCCATTT TCAAGAAAGC ATTGCCAGCT CTTCCAATCT
1561 CACTACCTT TGGGCTTGTG TTCTACTTTG CCACAGATTA TCTTGACAG CTTTTATGG
1621 ACCAATTAGC ATTCCATCAA TTTTATATCT AGCATATTTG CGGTAGAAAT CCCATGGATG
1681 TTTCTTCTTT GACTATAACA AAATCTGGGG AGGACAAAGG TGATTTTCTT GTGTCCACAT
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1801 CCTTGCACTA TTGGACTTTG GAAGGAGGTG CCTATAGAAA ACGATTTTGA ACATACTTCA
1861 TCGCAGTGGG CTGTGTCCCT CGGTGCAGAA ACTACCAGAT TTGAGGGACG AGGTCAAGGA
1921 GATATGATAG GCCCGAAGT TGCTGTGCCC CATCAGCAGC TTGACGCGTG GTCACAGGAC

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1981 GATTTCACTG ACACTGCGAA CTCTCAGGAC TACCGTTACC AAGAGGTTAG GTGAAGTGGT
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2101 TATTAAGTGA ATTCTGAACT TTTCAGGAGG TACTGTGAGG AAGAGCAGGC ACCAGCAGCA
2161 GAATGGGGAA TGGAGAGGTG GGCAGGGGTT CCAGCTTCCC TTTGATTTTT TGCTGCAGAC
2221 TCATCCTTTT TAAATGAGAC TTGTTTTCCC CTCTCTTTGA GTCAAGTCAA ATATGTAGAT
2281 TGCCTTTGGC AATTCTTCTT CTCAAGCACT GACACTCATT ACCGTCTGTG ATTGCCATTT
2341 CTTCCCAAGG CCAGTCTGAA CCTGAGGTTG CTTTATCCTA AAAGTTTTAA CCTCAGGTTT
2401 CAAATTCAGT AAATTTTGGA AACAGTACAG CTATTTCTCA TCAATTCTCT ATCATGTTGA
2461 AGTCAAATTT GGATTTTCCA CCAAATTCTG AATTTGTAGA CATACTTGTA CGCTCACTTG
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2581 CAGTAAGGCA GCTCTGTCTG GGTAGCAGAT GGTCCCATTA TTCTAGGGTC TTAATCTTTG
2641 TATGATGAAA AGAATGTGTT ATGAATCGGT GCTGTCAGCC CTGCTGTGAG ACCTTCTTCC
2701 ACAGCAAATG AGATGTATGC CCAAAGACGG TAGAATTAAA GAAGAGTAAA ATGGCTGTTG
2761 AAGC

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GENBANK ID: XM_087242.1
DEFINITION HOMO SAPIENS ARGINYL AMINOPEPTIDASE (AMINOPEPTIDASE B)-LIKE
1(RNPEPL1), MRNA.
VERSION XM_087242.1 GI:18600482
CDS 700..1764
/CODON_START=1

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121 TGGCTGGACC CAGAGCTGAC CTATGGCTGC GCCAAGCCCT TCGTCTTCAC CCAGGGCCAC
181 TCCGTGTGCA ACCGCTCCTT CTTCCTGTC TTCGACACAC CTGCCGTGAA GTGCACCTAC
241 TCTGCCGTCG TCAAGGCGCC ATCGGGGGTG CAGGTGCTGA TGAGTGCCAC CCGGAGTGCA
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961 CCACAGCGCT TTGATGACTT TCTCCGAGCC TATGTGGAGA AGTACAAGTT CACCAGCGTG
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GENBANK ID: S90469
DEFINITION CYTOCHROME P450 REDUCTASE [HUMAN, PLACENTA, MRNA PARTIAL, 2403 NT].
VERSION S90469.1 GI:247306
CDS 1..2031
/CODON_START=1

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181 TCCTCTGTCA GAGAGAGCAG CTTTGTGGAA AAGATGAAGA AAACGGGGAG GAACATCATC
241 GTGTCTACG GCTCCCAGAC GGGGACTGCA GAGGAGTTTG CCAACCGCCT GTCCAAGGAC
301 GCCCACCCTG ACGGGATGCG AGGCATGTCA GCGGACCCTG AGGAGTATGA CCTGGCCGAC
361 CTGAGCAGCC TGCCAGAGAT CGACAACGCC CTGTTGGTTT TCTGCATGGC CACCTACGGT
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1021 CTGGACGTCG TCATGTCCCT GAACAACCTG GATGAGGAGT CCAACAAGAA GCACCCATTTC
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1981 ATCAAGAAAC TGATGACCAA GGGCCGCTAC TCCCTGGACG TGTGGAGCTA GGGGCTGCC
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2161 TGGGCTGGGG GTGCATCCTC CTCAGCCCCC AGGCCAGGTG AGGTCCACCG GCGGCTGGCA
2221 GCACAGCCCA GGGCCTGCAT GGGGGCACCG GGCTCCATGC CTCTGGAGCC TCTGGCCCTC
2281 GGTGGCTGCA CAGAAGGGCT CTTTCTCTCT GCTGAGCTGG CCCAGCCCCC CCACGTGATT
2341 TCCAGTGAGT GTAAATAATT TTAATAAACC TCTGGCCCTT GGAATAAAGT TCTGTTTCT
2401 GTA
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GENBANK ID: NM_006254.1
DEFINITION HOMO SAPIENS PROTEIN KINASE C, DELTA (PRKCD), MRNA.
VERSION NM_006254.1 GI:5453969
CDS 59..2089
/CODON_START=1

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361 GGCTGAGTTC TGGCTGGACC TGCAGCCTCA GGCCAAGGTG TTGATGCTCT TTCAGTATT
421 CCTGGAGGAC GTGGATTGCA AACAATCTAT GCGCAGTGAG GACGAGGCCA AGTTCCCAAC
481 GATGAACCGC CGCGGAGCCA TCAAAACAGC CAAAATCCAC TACATCAAGA ACCATGAGTT
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661 CGACAAGATC ATCGGCAGAT GCACTGGCAC CGCGGCCAAC AGCCGGGACA CTATATTCCA
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781 CACCTTCTGT GACCACTGCG GCAGCCTGCT CTGGGGACTG GTGAAGCAGG GATTAAAGTG
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961 ATCAGACTCA GCCTCCTCAG AGCCTGTTGG GATATATCAG GGTTTCGAGA AGAAGACCCG
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1381 CAAAGGCCGC TTTGAACTCT ACCGTGCCAC GTTTTATGCC GCTGAGATAA TGTGTGGACT
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1981 GGGCGCGCTC TCCTACAGCG ACAAGAACCT CATCGACTCC ATGGACCAGT CTGCATTGCG
2041 TGGCTTCTCC TTTGTGAACC CCAAATTCGA GCACCTCTCG GAAGATTGAG GTTCTGGAC
2101 AGAT

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GENBANK ID: X61971.1

DEFINITION H.SAPIENS MRNA FOR MACROPAIN SUBUNIT DELTA.

VERSION X61971.1 GI:296733

CDS <1..543

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121 CACAGCATTG AACTGAATGA GCCTCCACTG GTCCACACAG CAGCCAGCCT CTTAAGGAG
181 ATGTGTTACC GATACCGGGA AGACCTGATG GCGGGAATCA TCATCGCAGG CTGGGACCTT
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301 GCCATTGGAG GCTCCGGGAG CTCCTACATC TATGGCTATG TTGATGCTAC CTACCGGGAA
361 GGCATGACCA AGGAAGAGTG TCTGCAATTC ACGGCCAATG CTCTCGCTTT GGCCATGGAG
421 CGGGATGGCT CCACTGGAGG AGTGATCCGC CTGGCAGCCA TTGCAGAGTC AGGGGTAGAG
481 CGGCAAGTAC TTTTGGGAGA CCAGATACCC AAATTCGCCG TTGCCACTTT ACCACCGGCC
541 TGAATCCTGG GATTCTAGTA TGCAATAAGA GATGCCCTGT ACTGATGCAA AATTTAATAA
601 AGTTTGTGAC AGAGAAAAAA AAAA

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45 GENBANK ID: AH005909.1

GENBANK ID: XM 088424.1

DEFINITION HOMO SAPIENS RETINOID X RECEPTOR, ALPHA (RXRA), MRNA.

VERSION XM_088424.1 GI:18571706

CDS 519..1016

/CODON_START=1

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1 AAGCAGAACG TGGCTCCCTT GGCCACAGCA GCCTTACCCA CCGCTCTACG TGTCCCGGGC
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241 TTATTATTTT GGTGGGACAA TCTTTAATTT TCTAAAGATA GCACTAACAT CAGCTCATTA
301 GCCACCTGTG CCTGTCCCGG CCTTGGCCCG GCTGGATGAA GCGGCTTCCC CGCAGGGCCC
361 CCACCTCCCA GTGGCTGCTT CCTGGGGACC CAGGGCACCC CGGCACCTTC AGGCACGCTC
421 CTCAGCTGGT CACCTCCCGG CTTTGCCGTT CAGATGGGGC TCCTGAGGCT CAGGAGTGAA
481 GATGCCACAG AGCCGGGCTC CCCTAGGCTG CGTCGGGCAT GCTTGAAGC TGGCCTGCCA
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661 GGGCTTGGCG CCTGGTGGCA GTGCTGGAGA TGACCCCGAG CCCCTCCCGG TGGGGCACCC
721 AGGAGGGCCC TGCCGAAATG TGCAGCTGTG GGGTAGTCGG CTGGTGTCCC TGTCTGGAG
781 CTGGGGTGCG TGATCTGCTG CTCGTCCACG CAGGTGTGTG GTGTAAACAT GTATGTGCTG

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841 TACAGAGAGA CGCGTGTGGA GAGAGCCGCA CACCAGCGCC ACCCAGGAAA GGCGGAGCGG
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961 CTTTCGTGTAA GCAAGTACAT AAGGACCCTC CTTTGGTGAA ATCCGGGTTC GAATGAATAT
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1381 ACCGCCGCAA TGGGGGTGTC TTCCCTCGGG GCAGGAGGGT GGGCCTGAGG CTTTCAAGGG
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1561 GCCGAGGTG CCAGGGTTTG ATGGACAGTA GCATTAGAAT TGTGGAAGG GAACACGCAA
1621 AGGGAGAAGT GTGAGAGGAG AAACAAAATA TGAGCGTTTA AAATACATCG CCATTTCAG

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GENBANK ID: U41745.1
DEFINITION HUMAN PDGF ASSOCIATED PROTEIN MRNA, COMPLETE CDS.
VERSION U41745.1 GI:1136583
CDS 22..567
/CODON_START=1

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1 GAATTCCGCG GCGGCGCCTC AATGCCTAAA GGAGGAAGAA AGGGAGGCCA CAAAGGCCGG
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121 GCCAGGGAAG AAGAGGAGCA AAAAGAAGGT GGAGATGGGG CTGCAGGTGA CCCCCAAAAG
181 GAGAAGAAAT CTCTAGACTC AGATGAGAGT GAGGATGAAG AAGATGACTA CCAGCAAAAG
241 CGCAAAGGCG TTGAAGGGCT CATCGACATC GAGAACCCCA ACCGGGTGGC ACAGACAACC
301 AAAAAGGTCA CACAACCTGA TCTGGACGGG CCAAAGGAGC TTTCGAGGAG AGAACGAGAA
361 GAGATTGAGA AGCAGAAGGC AAAAGAGCGT TACATGAAAA TGCACTTGGC CGGGAAGACA
421 GAGCAAGCCA AGGCTGACCT GGCCCGGCTG GCCATCATCC GGAACACAGC GGAGGAGGCT
481 GCCCGGAAGA AGGAAGAGGA AAGGAAGCA AAAGACGATG CCACATTGTC AGGAAACGTA
541 ATGCAGTCAC TCTCCCTGAA TAAGTAACTG CGACCCGTGG GAGGAGATGC CGGGGACCTG
601 GGCCGCGCTG CCAGGACCTC TGCTGTGTCT CGCCACCCCT GTGCCCTGGC GCCGCTGCAA
661 CAGCCCTCA TGGCCAGGAG CCCCCATGC CTGGGCTTCC TCTTCATCTT GGCACAGAAA
721 TTGTTTGGGG GATGGGGGGG GGAAGTGGGG AGGGGTAGCT GCTATCTTTG AGACAGAAAG
781 ATGCAGGACA GCATTTCATA TGAACCAATT TGAATGTTTT TGCTGTTTTT AGAATTTC

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GENBANK ID: AH002617.1

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GENBANK ID: XM_034862.1
DEFINITION HOMO SAPIENS INTERFERON REGULATORY FACTOR 1 (IRF1), MRNA.
VERSION XM_034862.1 GI:14726087
CDS 197..1174
/CODON_START=1

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1 CGAGCCCCGC CGAACCAGAG CCACCCGGAG CCGTGCCCAG TCCACGCCGG CCGTGCCCCG
61 CGGCCCTTAAG AACCCGGCAA CCTCTGCCTT CTTCCCTCTT CCACTCGGAG TCGCGCTCCG
121 CGCGCCCTCA CTGCAGCCCC TGCGTCGCGG GGACCTCGC GCGCGACCGC CGAATCGCTC
181 CTGCAGCAGA GCCAACATGC CCATCACTCG GATGCGCATG AGACCCCTGGC TAGAGATGCA
241 GATTAATTCC AACCAAATCC CGGGGCTCAT CTGGATTAAT AAAGAGGAGA TGATCTTCCA
301 GATCCCATGG AAGCATGCTG CCAAGCATGG CTGGGACATC AACCAAGGATG CCTGTTTGT
361 CCGGAGCTGG GCCATTACA CAGGCCGATA CAAAGCAGGG GAAAAGGAGC CAGATCCCAA
421 GACGTGGAAG GCCAATTTT GCTGTGCCAT GAACTCCCTG CCAGATATCG AGGAGGTGAA
481 AGACCAGAGC AGGAACAAGG GCAGCTCAGC TGTGCGAGTG TACCGGATGC TTCCACCTCT
541 CACCAAGAAC CAGAGAAAAG AAAGAAAGTC GAAGTCCAGC CGAGATGCTA AGAGCAAGGC
601 CAAGAGGAAG TCATGTGGGG ATTCCAGCCC TGATACCTTC TCTGATGGAC TCAGCAGCTC
661 CACTCTGCCT GATGACCACA GCAGCTACAC AGTTCCAGGC TACATGCAGG ACTTGGAGGT
721 GGAGCAGGCC CTGACTCCAG CACTGTGCGC ATGTGCTGTC AGCAGCACTC TCCCGACTG
781 GCACATCCCA GTGGAAGTTG TGCCGGACAG CACCAGTGAT CTGTACAACT TCCAGGTGTC
841 ACCCATGCCC TCCACCTCTG AAGCTACAAC AGATGAGGAT GAGGAAGGGA AATTACCTGA
901 GGACATCATG AAGCTCTTGG AGCAGTCGGA GTGGCAGCCA ACAACGCTGG ATGGGAAGGG
961 GTACCTACTC AATGAACCTG GAGTCCAGCC CACCTCTGTC TATGGAGACT TTAGCTGTAA
1021 GGAGGAGCCA GAAATGACA GCCCAGGGGG GGATATTGGG CTGAGTCTAC AGCGTGTCTT
1081 CACAGATCTG AAGAACATGG ATGCCACCTG GCTGGACAGC CTGCTGACCC CAGTCCGGTT
1141 GCCCTCCATC CAGGCCATTG CCTGTGCACC GTAGCAGGGC CCCTGGGCCC CTCTTATTC

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1201 TCTAGGCAAG CAGGACCTGG CATCATGGTG GATATGGTGC AGAGAAGCTG GACTTCTGTG
1261 GGGCCCTCAA CAGCCAAGTG TGACCCCACT GCCAAGTGGG GATGGGGCCT CCCTCCTTGG
1321 GTCATTGACC TCTCAGGGCC TGGCAGGCCA GTGTCTGGGT TTTTCTTGTG GTGTAAGCT
1381 GGGCCTGCCT CCTGGGAAGA TGAGGTCTGT AGACCAGTGT ATCAGGTGAG GGACTTGGAC
1441 AGGAGTCAGT GTCTGGCTTT TTCCTCTGAG CCCAGCTGCC TGGAGAGGGT CTCGCTGTCA
1501 CTGGCTGGCT CCTAGGGGAA CAGACCAGTG ACCCCAGAAA AGCATAACAC CAATCCCAGG
1561 GCTGGCTCTG CACTAAGAGA AAATTGCACT AAATGAATCT CGTTCCCAA GAACATCCCC
1621 CTTTTCAGCT GAGCCCTGGG GACTGTTCCA AAGCCAGTGA AATGTGAAGG AAAGTGGGGT
1681 CCTTCGGGGC GATGCTCCCT CAGCCTCAGA GGAGCTCTAC CCTGCTCCCT GCTTTGGCTG
1741 AGGGGCTTGG GAAAAAACT TGGCACTTTT TCGTGTGGAT CTGCCCACAT TTCTGATCAG
1801 AGGTGTACAC TAACATTTC CCCGAGCTCT TGGCCTTTGC ATTTATTTAT ACAGTGCCTT
1861 GCTCGGCGCC CACCACCCCC TCAAGCCCCA GCAGCCCTCA ACAGGCCAGG GGAGGGAGT
1921 GTGAGCGCCT TGGTATGACT TAAAATTGGA AATGTCATCT AACCATTAAG TCATGTGTGA
1981 ACACATAGGA CGTGTGTAAA TATGTACATT TGTCTTTTAA TAAAAAGTAA ATTGTT

GENBANK ID: AJ310549.1

DEFINITION HOMO SAPIENS MRNA FOR CLP-36 PROTEIN.

VERSION AJ310549.1 GI:13160404

CDS 1..990

/CODON_START=1

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1 ATGACCACCC AGCAGATAGA CCTCCAGGGC CCGGGGCCGT GGGGCTTCCG CCTCGTGGGC
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121 CTAGCTAATT TATGTATTGG AGATGTAATC ACAGCCATTG ATGGGGAAAA TACTAGCAAT
181 ATGACACACT TGGAGCTCA GAACAGAATC AAAGCTGCA CAGACAACTT GACTCTCACT
241 GTAGCCAGAT CTGAACATAA AGTCTGTGCT CCTCTGGTGA CGGAGGAAGG GAAGCGTCAT
301 CCATACAAGA TGAATTTAGC CTCTGAACCC CAGGAGGTCC TGCACATAGG AAGCGCCAC
361 AACCGAAGTG CCAATGCCCTT TACCGCCTCG CCTGCCTCCA GCCTACTGTC CAGGGTCATC
421 ACAAAACAGT ACAACAACCC AGCTGGCCTC TACTCTTCTG AAAATATCTC CAACTTCAAC
481 AATGCCCTGG AGTCAAAGAC TGCTGCCAGC GGGGTGGAGG CGAACAGCAG ACCCTTAGAC
541 CATGCTCAGC CTCCAAGCAG CCTGTGCATC GACAAAGAAT CTGAAGTTTA CAAGATGCTT
601 CAGGAGAAAC AGGAGTTGAA TGAGCCCCCG AAACAGTCCA CGTCTTTCTT GGTTTTGCAG
661 GAAATCCTGG AGTCTGAAGA AAAAGGGGAT CCCAACAAGC CCTCAGGATT CAGAAGTGT
721 AAAGCTCCTG TCACTAAAGT GGTGCGTCG ATTGGAAATG CTCAGAAGTT GCCTATGTGT
781 GACAAATGTG GCACTGGGAT TGTGGTGTG TTTGTGAAGC TGGGGGACCG TCACCGCCAC
841 CCTGAGTGTT ATGTGTGCAC TGACTGTGGC ACCAACCTGA AACAGAAGGG CCATTTCTTT
901 GTGGAGGATC AAATCTACTG TGAGAAGCAT GCCCGGGAGC GAGTCACACC ACCTGAGGGT
961 TATGAAGTGG TCACTGTGTT CCCCAAGTGA GCCAGCAGAT CTGACCACTG TTCTCCAGCA
1021 GGCCTCTGCT GCAGCTTTT CTCTCAGTGT TCTGGCCCTC TCCTCTCTTG AAAGTTCTCT
1081 GCTTACTTTG GTT

GENBANK ID: XM_016642.3

DEFINITION HOMO SAPIENS ADENYLATE KINASE 3 (AK3), MRNA.

VERSION XM_016642.3 GI:16163712

CDS 145..816

/CODON_START=1

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61 GGGCCGAGAA ACAAGTTCC CGGGGCTCCC TCCGGGGCCG CGGTGCGGGC TCGCGGTTG
121 ACCGCCCCCC TCCTCGCGAA GGCAATGGCT TCCAACTCC TGGCGCGGCT CATCCTCGGG
181 CCGCCCGGCT CGGGCAAGGG CACCGTGTGC CAGAGGATCG CCCAGAACTT TGGTCTCCAG
241 CATCTCTCCA GCGGCCACTT CTGCGGGAG AACATCAAGG CCAGCACCAG AGTTGGTGAG
301 GTGGCAAAGC AGTATATAGA GAAAAGTCTT TTGGTTCCAG ACCATGTGAT CACACGCCA
361 ATGATGTCCG AGTTGGAGAA TAGGCGTGGC CAGCACTGGC TCCTTGATGG TTTCTCTAGG
421 ACATTAGGAC AAGCCGAGGC CCTGGACAAA ATCTGTGAAG TGGATCTAGT GATCAGTTT
481 AATATTCCAT TTGAAACACT TAAAGATCGT CTCAGCCGCC GTTGGATTCA CCTCTCTAGC
541 GGAAGGGTAT ATAACCTGGA CTTCATCCA CCTCATGTAC ATGGTATTGA TGACGTCACT
601 GGTGAACCAT TAGTCCAGCA GGAGGATGAT AAACCGAAG CAGTTGCTGC CAGGCTAAGA
661 CAGTACAAAG ACGCGGCAAA GCCAGTCATT GAATTATACA AGAGCCGAGG AGTGCTCCAC
721 CAATTTTCCG GAACGGAGAC GAACAAATC TGGCCCTACG TTTACACACT TTTCTCAAAC
781 AAGATCACAC CTATTCAGTC CAAAGAAGCA TATTGACCCT GCCCAATGGA AGAACCAGGA
841 AGATGTGGTC ATTCAATCAA TAGTGTGTGT AGTATTGGTG CTGTGTCCAA ATTAGAAGCT
901 AGCTGAGGTA GCTTGACGCA TCTTTCTAG TTGAAATGGT GAACTGATAG GAAACAAAT
961 GAGTAGAAG AGTTCATGAA GAGGCCCTCC TCTGCCTTC AAAAGGGTGG TCACCTACAC
1021 ATGTTTAAGG TGTCTCTGCA CATGTCTCAA GCCCATCACA AGAAGCAAG TACAGTGTGG

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1081 ATTTCAAATG GTGTGTAAC TCAAGCTCCAG CTGGTTTTTG ACAGCTGTTG CTGTGGTAAT
1141 ATTTTTTACA TGTGATGGTG ATAGTCTCTG GTTCTCCCCA TCCCCACAAA GGCTGTTGAA
1201 CCACAGCACC AGGAAGCCTG AGAATGAATC CTGAGGGCTC TAGCCCAGGC TTGTGCCCAG
1261 GCTTTCTGGT GTGTGCCCTC CTGGTAACAG TGAAATTGAA GCTACTTACT CATAGTGGTT
1321 GTTTCTCTGG TCTTGAGTGA CTGTGTCCAC AGTTCATTTT TTCCCGGTAG GAATAACTCC
1381 TTTTCTACAT CCACACTCCA TAGAGTCTCT CCTTTTCAGA TATCCTGGGA TGAAAGAATT
1441 TGGCTTTTTT TTTTTTTTTT TTTTGGACAT CTGTTTTCAC TCTTAGGCTT TTAACAATA
1501 GTTATTGCTC TTATCCCTCT CAGATTCTAA TAACTGAGAG TGATGGGGCT ATATTGAATC
1561 TCTGTATGCA CTGAGAACTG AGCTATGAAG AGGATCTTAT TAAACTGCTG GTCTGACTTT
1621 ATGGATTGAC ACTGTTTCCT TCTTTTATTG TG

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GENBANK ID: Z35307.1
DEFINITION H.SAPIENS MRNA FOR ENDOTHELIN-CONVERTING-ENZYME 1.
VERSION 235307.1 GI:535181
CDS 38..2299
/CODON_START=1

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1 CGCCCCCCCG GTGTCCGCC TGCTGTCCGG GCTGGGGATG TCGACGTACA AGCGGGCCAC
61 GCTGGACGAG GAGGACCTGG TGGACTCGCT CTCCGAGGGC GACGCATACC CCAACGGCCT
121 GCAGGTGAAC TTCCACAGCC CCCGGAGTGG CCAGAGGTGC TGGGCTGCAC GGACCAGGT
181 GGAGAAGCGG CTGGTGGTGT TGGTGGTACT TCTGGCGGCA GGACTGGTGG CCTGCTTGGC
241 AGCACTGGGC ATCCAGTACC AGACAAGATC CCCCTCTGTG TGCCTGAGCG AAGCTTGTGT
301 CTCAGTGACC AGCTCCATCT TGAGCTCCAT GGACCCACA GTGGACCCCT GCCATGACTT
361 CTTAGCTAC GCCTGTGGGG CTGGGATCAA GGCCAAACCA GTCCCTGATG GCCACTCAGC
421 CTGGGGGACC TTCAGCAACC TCTGGGAACA CAACCAAGCA ATCATCAAGC ACCTCTCGA
481 AAATCCACG GCCAGCGTGA GCGAGGCAGA GAGAAAGGCG CAAGTATACT ACCGTGCGTG
541 CATGAACGAG ACCAGGATCG AGGAGCTCAG GGCCAAACCT CTAATGGAGT TGATTGAGAG
601 GCTCGGGGGC TGGACATCA CAGGTCCCTG GGCCAAGGAC AACTTCCAGG ACACCTGCA
661 GGTGGTCACC GCCCACTACC GCACCTCACC CTTCTTCTCT GTCTATGTCA GTGCCGATT
721 CAAGAATCC AACAGCAACG TGATCCAGGT GGACCAGTCT GGCCCTGGGCT TGCCCTCGAG
781 AGACTATTAC CTGAACAAAA CTGAAAACGA GAAGGTGCTG ACCGGATATC TGAATACAT
841 GGTCCAGCTG GGGAGCTGC TGGGCGGCGG GGACGAGGAG GCCATCCGGC CCCAGATGCA
901 GCAGATCTTG GACTTTGAGA CGGCCTGGC CAACATCACC ATCCCACAGG AGAAGCGCCG
961 TGATGAGGAG CTCATCTACC ACAAGTGAC GGCAGCCGAG CTGCAGACCT TGGCACCCTG
1021 CATCAACTGG TTGCCCTTTT TCAACACCAT CTTCTACCCC GTGGAGATCA ATGAATCCGA
1081 GCCTATTGTG GTCTATGACA AGGAATACCT TGAGCAGATC TCCACTCTCA TCAACACCAC
1141 CGACAGATGC CTGCTCAACA ACTACATGAT CTGGAACCTG GTGCGGAAAA CAAGCTCCTT
1201 CCTTGACCAG CGCTTTTCAGG ACGCCGATGA GAAGTTTATG GAAGTCATGT ACGGGACCAA
1261 GAAGACCTGT CTTCTCGCT GGAAGTTTGT CGTGAGTGAC ACAGAAAACA ACCTGGGCTT
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1381 CGAGATCATC CTGGAGATTA AGAAGGCATT TGAGGAAAGC CTGAGCACCC TGAAGTGGAT
1441 GGATGAGGAA ACCCGAAAAT CAGCCAAGGA AAAGGCCGAT GCCATCTACA ACATGATAGG
1501 ATACCCCAAC TTCATCATGG ATCCCAAGGA GCTGGACAAA GTGTTTAATG ACTACACTGC
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1621 TGCCGATCAG CTCAGGAAAG CCCCCAACAG AGATCAGTGG AGCATGACCC CGCCCATGGT
1681 GAACGCCTAC TACTCGCCCA CCAAGAATGA GATTGTGTTT CCGGCCGGGA TCCTGCAGGC
1741 ACCATTCTAC ACACGCTCCT CACCAAGGC CTTAACTTTT GGTGGCATAG GTGTCGTCGT
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1861 CCTCCGGCCA TGGTGGAAAG ACTCATCCGT GGAGGCCTTC AAGCGTCAGA CCGAGTGCAT
1921 GGTAGAGCAG TACAGCAACT ACAGCGTGAA CGGGGAGCCG GTGAACGGGC GGCACACCCT
1981 GGGGGAGAAC ATCGCCGACA ACGGGGGTCT CAAGGCGGCC TATCGGGCTT ACCAGAACTG
2041 GGTGAAGAAG AACGGGGCTG AGCACTCGCT CCCACCCCTG GGCCTCACCA ATAACAGCT
2101 CTTCTTCCTG GGCTTTGCAC AGGTCTGGTG CTCCGTCCGC ACACCTGAGA GCTCCCACGA
2161 AGGCCTCATC ACCGATCCCC ACAGCCCCCTC TCGCTTCGGG GTCATCGGCT CCCTCTCAA
2221 TTCCAAGGAG TTCTCAGAAC ACTTCCGCTG CCCACCTGGC TCACCCATGA ACCCGCTCA
2281 CAAGTGCGAA GTCTGGTAAG GACGAAGCGG AGAGAGCCAA GACGGAGGAG GGAAGGGGC
2341 TGAGGACGAG ACCCCCATCC AGCCTECAGG GCATTGCTCA GCCCGCTTGG CCACCCGGG
2401 CCCTGCTTCC TCACACTGGC GGGTTTTTCT CCGGAACCGA GCCCATGGTG TTGGCTCTCA
2461 ACGTGACCCG CAGTCTGATC CCTGTGAAG AGCCGGACAT CCCAGGCACA CGTGTGCGCC
2521 ACCTTCAGCA GGCATTGGGG GTGTGGGCTG GTGGCTCATC AGGCCTGGGC CCCACTGA
2581 CAAGCGCCAG ATACGCCACA AATACCCTG TGTCAAATGC TTTCAAGATA TATTTTGGG
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2701 ACACTTTTTT TTTTAAGCCC

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GENBANK ID: J02683
DEFINITION HUMAN ADP/ATP CARRIER PROTEIN MRNA, COMPLETE CDS.

VERSION J02683.1 GI:179246
CDS 70..966
/CODON_START=1

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121 GCCGCAGCCA TCTCCAAGAC GGCGGTAGCG CCCATCGAGC GGGTCAAGCT GCTGCTGCAG
181 GTGCAGCATG CCAGCAAGCA GATCACTGCA GATAAGCAAT ACAAAGGCAT TATAGACTGC
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10 301 GTCATCAGAT ACTTCCCCAC CCAGGCTCTT AACTTCGCCT TCAAAGATAA ATACAAGCAG
361 ATCTTCCTGG GTGGTGTGGA CAAGAGAACC CAGTTTTGGC GCTACTTTGC AGGGAATCTG
421 GCATCGGGTG GTGCCGCGAG GGCCACATCC CTGTGTTTG TGTACCCTCT TGATTTTGCC
481 CGTACCCGTC TAGCAGCTGA TGTGGGTAAA GCTGGAGCTG AAAGGGAATT CCGAGGCCCTC
541 GGTGACTGCC TGGTTAAGAT CTACAAATCT GATGGGATTA AGGGCCTGTA CCAAGGCTTT
15 601 AACGTGCTG TGCAGGGTAT TATCATCTAC CGAGCCGCCT ACTTCGGTAT CTATGACACT
661 GCAAAGGGAA TGCTTCCGGA TCCCAAGAAC ACTCACATCG TCATCAGCTG GATGATCGCA
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781 ATGATGATGC AGTCAGGGCG CAAAGGAAC GACATCATGT ACACAGGCAC GCTTGACTGC
841 TGGCGGAAGA TTGCTCGTGA TGAAGGAGGC AAAGCTTTT TCAAGGGTGC ATGGTCCAAT
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1021 AATCTTGAGC ATTCTTGACA GACTCCTGGC TGTCAAGTTT TCAGTGGCAA CTACTTTACT
1081 GGTGAAAAAT GGAAGCAAT AATATTCATC TGACCAGTTT TCTCTTAAAG CCATTTCCAT
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25 1201 ATTTGGAGAA ATAAAAATAG TCTAAAA

GENBANK ID: M22760.1
DEFINITION HOMO SAPIENS NUCLEAR-ENCODED MITOCHONDRIAL CYTOCHROME C OXIDASE VA
SUBUNIT MRNA, COMPLETE CDS.

30 VERSION M22760.1 GI:695359
20..472
/CODON_START=1

35 1 GGGCGCCGCC ATCGCCGTCA TGCTGGGCGC CGCTCTCCGC CGCTGCGCTG TGGCCGCAAC
61 CACCCGGGCC GACCCCTCGAG GCCTCCTGCA CTCCGCCCGG ACCCCCGGCC CCGCCGTGGC
121 TATCCAGTCA GTTCGCTGCT ATTCCTCATGG GTCACAGGAG ACAGATGAGG AGTTTGATGC
181 TCCTGGGTA ACATACTTCA ACAAGCCAGA TATAGATGCC TGGGAATTGC GTAAAGGGAT
241 AAACACACTT GTTACCTATG ATATGGTTCC AGAGCCCAA ATCATTGATG CTGCTTTGCG
40 301 GGCATGCAGA CGGTTAAATG ATTTTGCTAG TCTAGTTCGA ATCTAGAGG TTGTTAAGGA
361 CAAAGCAGGA CCTCATAAGG AAATCTACCC CTATGTCTAT CAGGAACCTA GACCAACTTT
421 AATGAACCTG GGAATCTCCA CTCGGGAGGA ACTGGGCCTT GACAAAGTGT AAACCGCATG
481 GATGGGCTTC CCAAGGATT TATTGACATT GCTACTTGAG TGTGAACAGT TACCTGGAAA
541 TACTGATGAT AACATATTAC CTTATTTTGA ACAAGTTTCC CTTTATTGAG TACCAAGCCA
45 601 TGTAATGGTA ACTTGACTT TAATAAAGG GAAATGAGTT TGAAGT

GENBANK ID: M18079
DNA LINEAR
DEFINITION HUMAN, INTESTINAL FATTY ACID BINDING PROTEIN GENE, COMPLETE CDS, AND
AN ALU REPETITIVE ELEMENT.

50 VERSION M18079.1 GI:182351

1 GTAATATCTT GGGCAAGCCC TAGAGCTTCT TTCCTGACCC TTAGTTAATA AGATGTTATC
61 TGGTCACATT CAGTCACAAT AATAGACTCA TTTTAGTAAT AAACATCTTA AGACTAGTAA
55 121 TTAAAACTCT TTAATTCACA CCAAGTTTCC TCCCCAAGCT TGGCCTGTTT CTGGCTGGCA
181 GCCTGAAGTA GGGAAAGGAG AGATATGGTG ACCTTTTCTT TGTACCTTTC TAGCTACCTT
241 CTATACCCTG ACCCCACATA CATAATTGAG CTGTGGCTTC TGACTCTACT GGGTTTGGGG
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421 TTTACACCAT CCAGCCTCAT TTGTACCTCT TGAATTTTGT CTCAGTGGCC TATCACCATT
60 481 CAGGATCAAG ACAAAAATCA ATGAGCACTT ATTGTGTGTC ATGCACCTTA CAAAGTGCCA
541 GGATATTTAT CCAAACTCCT GGCAATGCTA AACACAATGC AAAAAGACAT ATTAGAAAAC
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661 AATTGCTGTT GTGAAAAAGG GAAAGTCAT GGTCTCATTT CCCAGATGTT ATTTGATATA
721 TGCTATAAAT TATATTACCT CCAACATAGT CTGCACCTTG AACTTAGAAA AACATCTTC
65 781 AGACGGCATG CATTCTAATT CTTGAAATAA GTATGCCAC AAAGTGTAGT TTAAGACAGA
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5 901 AACGAGAATT AAGAATTAAT AAGAATAAGA ATTAATTAAT TGCTTGACAT AGAGTAGTTA
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 2281 ATAACCTTGA AAAATAAACA CTTCTATGG GATTTGACTT TATTTTCTCC ATTGTCTTAC
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 2641 GAAGCCAATG AAGTTTGTCT ACATTATATA TTACACAAAT TGGCAGGTA TTTAAATATG
 2701 CTTTATTTT TATACGCATC TGTGAAGAAT CTGAATTGAA CAGTAAGAAAT TAGAAAATA
 2761 TCTTTTGAAT GACTGAATAT AGACCTATTC ATAAAGAAAT TTTAACTGT GTTTTAAAC
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 2881 TAAAAGAAAT GAATAGATGA ACAAATGAGT GAGTTACCAA ATGGAAGAT TTGATGTATT
 2941 GTAGTCAATT GGGAGTGTAC CTTTTCATGT TTAAGATAAC ACATTTTAGG AAGTCATCAT
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GENBANK ID: X16277

DNA LINEAR

DEFINITION HUMAN GENE FOR ORNITHINE DECARBOXYLASE ODC (EC 4.1.1.17).

VERSION X16277.1 GI:35137

MRNA JOIN(795..1001,3858..3967,4073..4191,4475..4648,

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DEFINITION H.SAPIENS RING4 CDNA.
VERSION X57522.1 GI:36060
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GENBANK ID: D00017.1
DEFINITION HOMO SAPIENS MRNA FOR LIPOCORTIN II, COMPLETE CDS.
VERSION D00017.1 GI:219909
CDS 50..1069
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GENBANK ID: M10277.1
DNA LINEAR
DEFINITION HUMAN CYTOPLASMIC BETA-ACTIN GENE, COMPLETE CDS.
VERSION M10277.1 GI:177967

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GENBANK ID: XM_042788.1

DEFINITION HOMO SAPIENS ALDOLASE B, FRUCTOSE-BISPHOSPHATE (ALDOB), MRNA.

VERSION XM_042788.1 GI:14738248

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 VERSION NM_005317.2 GI:7108347
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 CDS 1401..2957
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GENBANK ID: AF035429.1

DNA LINEAR

15 DEFINITION HOMO SAPIENS CYTOCHROME OXIDASE SUBUNIT I (COI) AND SUBUNIT II
(COII) PSEUDOGENES, COMPLETE SEQUENCE.
VERSION AF035429.1 GI:2665724

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45 1501 AATAATTTTC ATAATTGAG AAGCCTTCGC TTGGAAGCGA AAAGTCTTAA TAGTAGAAGA
1561 ACCCTCCATA AACCTGGAGT GACTATATGG ATGCCCCCA CCCTACCACA CATTCGAAGA
1621 ACCCGTATAC ATAAAATCTA GACAAAAAAG GAAGGAATCG AACCCCCAA AGCTGGTTTC
1681 AAGCCAACCC CATGGCCTCC ATGACTTTTT CAAAAAGATA TTAGAAAAAC CATTTCTATA
1741 CTTTGTCAAA GTTAAATTAT AGGCTAAATC CTATATATCT TAATGGCACA TGCAGCGCAA
50 1801 GTAGGTCTAC AAAACGCTAC TTCCCCTATC ATAGAAGAGC TTATCATCTT TCATGATCAC
1861 GCCCTCATAA TCATTTTCCT TATCTGCTTC CTAGTCTGT ACGCCCTTTT CCTAACACTC
1921 ACAACAAAC TAACTAATAC TAACATCTCA GACGCTCAGG AAATAGAAAC CGTCTGAAC
1981 ATCCTGCCCG CCATCATCCT AGTCCTTATC GCCCTCCCAT CCCTACGCAT CCTTTACATA
2041 ACAGACGAGG TCAACGATCC CTCCTTTACC ATCAAATCAA TTGGCCATCA ATGGTACTGA
55 2101 ACCTACGAAT ACACCGACTA CGGCGGACTA ATCTTCAACT CCTACATACT TCCCCATTA
2161 TTCCTAGAAC CAGGCGACCT GCGACTCCTT GACGTTGACA ATCGAGTAGT ACTCCCGGTT
2221 GAAGCCCCCA TTCGTATAAT AATTACATCA CAAGACGCTT TACACTCATG AGCTGTCCCC
2281 ACATTAGGCT TAAAAACAGA TGCAATTCCC GGACGCTTAA ACCAAACCAC TTTCACTGCT
2341 ACACGACGAG GGGTATACTA CGGCCAATGC TCTGAAATCT GTGGAGCAAA CCAGTTTAT
60 2401 GCCCATCGTC CTAGAATTAA TTCCCCTAAA AATCTTTGAA ATAGGCGCTG TATTTACCT
2461 ATAGCACCCC CTCTACCCCC TCTAGAGCCC ACTGTAAAGC TAACTTAGCA TTAACCTTTT
2521 AAGTTAAAGA TTAAGAGAAC CAACACCTCT TTACAGTGAA ATGCCCCAAC TAAATACTA

GENBANK ID: NM_024409.1

65 DEFINITION HOMO SAPIENS NATRIURETIC PEPTIDE PRECURSOR C (NPPC), MRNA.
VERSION NM_024409.1 GI:13249345

CDS 1..381
/CODON_START=1

5 1 ATGCATCTCT CCCAGCTGCT GGCCTGCGCC CTGCTGCTCA CGCTGCTCTC CCTCCGGCCC
61 TCCGAAGCCA AGCCCCGGGC GCCGCGAAG GTCCCGCGAA CCCC GCCGGC AGAGGAGCTG
121 GCCGAGCCGC AGGCTGCGGG CGGCGGTGAG AAGAAGGGCG ACAAGGCTCC CGGGGGCGGG
181 GCGCCCAATC TCAAGGGCGA CCGGTCGCGA CTGCTCCGGG ACCTGCGCGT GGACACCAAG
241 TCGCGGGCAG CGTGGGCTCG CCTTCTGCAA GAGCACCCA ACGCGCGCAA ATACAAAGGA
301 GCCAACAGA AGGGCTTGTC CAAGGGCTGC TTCGGCCTCA AGCTGGACCG AATCGGCTCC
10 361 ATGAGCGGCC TGGGATGTGA G

GENBANK ID: M37763.1
DNA LINEAR
DEFINITION HUMAN NEUROTROPHIN-3 (NT-3) GENE, COMPLETE CDS.
15 VERSION M37763.1 GI:189300
CDS 76..849
/CODON_START=1
GENE 76..849
20 MAT PEPTIDE 130..846

1 TAACACAGAC TCAGCTGCCA GAGCCTGCTC TTAACACCTG TGTTCCTTT TCAGATCTTA
61 CAGGTGAACA AGGTGATGTC CATCTGTGTT TATGTGATAT TTCTCGCTTA TCTCCGTGGC
121 ATCCAAGGTA ACAACATGGA TCAAAGGAGT TTGCCAGAAG ACTCGCTCAA TTCCCTCAT
181 ATTAAGCTGA TCCAGGCAGA TATTTGAAA AACAAGTCT CCAAGCAGAT GGTGGACGTT
25 241 AAGGAAAATT ACCAGAGCAC CCTGCCCAA GCTGAGGCTC CCGAGAGCC GGAGCGGGGA
301 GGGCCCGCCA AGTCAGCATT CCAGCCGGTG ATGCAATGG ACACCGAACT GCTGCGACAA
361 CAGAGACGCT ACAACTCACC GCGGGTCCTG CTGAGCGACA GCACCCCTT GGAGCCCCCG
421 CCCTTGTATC TCATGGAGGA TTACGTGGGC AGCCCCGTGG TGGCGAACAG AACATCACGG
481 CGGAAACGGT ACGCGGAGCA TAAGAGTCAC CGAGGGGAGT ACTCGGTATG TGACAGTGAG
30 541 AGTCTGTGGG TGACCGACAA GTCATCGGCC ATCGACATTC GGGGACACCA GGTACGGTG
601 CTGGGGGAGA TCAAAACGGG CAACTCTCCC GTCAAACAAT ATTTTATGA AACCGATGT
661 AAGGAAGCCA GGCCGGTCAA AAACGGTTC AGGGGTATTG ATGATAAACA CTGGAACCTC
721 CAGTGCAAAA CATCCCAAA CTACGTCCGA GCACTGACTT CAGAGAACA TAACTCGTG
781 GGCTGGCGGT GGATACGGAT AGACACGTCC TGTGTGTGTG CCTTGTGAG AAAAATCGGA
35 841 AGAACATGAA TTGGCATCTC TCCCATATA TAAATTATTA CTTTAAATTA TATGATATGC
901 ATGTAGCATA TAAATGTTTA TATTGTTTT ATATATTATA AGTTGACCTT TATTATTAA
961 ACTTCAGCAA CCCTACAGTA TATAAGCTTT TTTCTCAATA AAATCAGTGT GCTTGCCTTC

GENBANK ID: NM_000932.1
40 DEFINITION HOMO SAPIENS PHOSPHOLIPASE C, BETA 3
(PHOSPHATIDYLINOSITOL-SPECIFIC) (PLCB3), MRNA.
VERSION NM_000932.1 GI:11386138
CDS 1..3705

1 ATGGCGGGCG CCCAGCCCGG CGTCCACGCG CTGCAGTTGG AGCCGCCAC CGTGGTGGAG
61 ACCCTGCGGC GCGGGAGTAA GTTCATCAAA TGGGACGAGG AGACCTCCAG TCGBAACCTG
121 GTGACCTGCG GTGTGGACCC CAATGGCTTC TTCTTGTAAT GGACGGGCC CAACATGGAG
181 GTGACACAC TGGACATCAG TTCCATCAGG GACACACGGA CAGGCCGGTA CGCCCGCCTG
241 CCAAGGACC CCAAGATCCG GGAAGTTCTG GGCTTTGGGG GTCCCGATGC CCGGCTGGAG
50 301 GAGAAGCTGA TGACGGTGGT GTCTGGGCCA GACCCGGTGA ACACAGTGT CTTGAACCTC
361 ATGGCCGTGC AGGATGACAC AGCCAAGGTC TGGTCTGAGG AGCTATTCAG GCTGGCTATG
421 AACATCCTGG CTCAGAACGC CTCCCGGAAC ACCTTCCTGC GCAAAGCATA CACGAAGCTG
481 AAGCTGCAGG TGAACCAGGA TGGTCGGATC CCGTCAAGA ACATCCTGAA GATGTTCTCA
541 GCAGACAAGA AGCGGGTGGG GACTGCGCTG GAATCCTGTG GCCTCAAATT CAACCGGAGT
55 601 GAGTCCATCC GGCTGTATGA GTTTTCCTTG GAAATCTTTG AGCGGTTCCT GAACAAGCTG
661 TGTCTGCGGC CGGACATTGA CAAGATCCTG CTGGAGATAG GCGCCAAGGG CAAGCCATAC
721 CTGACGCTGG AGCAGCTCAT GGACTTCATC AACCAGAGGC AACCGGACCC GAGACTCAAC
781 GAAGTGCTGT ACCCGCCCTC GCGGCCCTCC CAGGCCCGGC TGCTCATCGA AAAGTATGAG
841 CCCAACACGC AGTTTCTGGA GCGAGACCAG ATGTCCATGG AGGGCTTTAG CCGTCACTG
60 901 GAGGGCGAGG AGAATGGCAT CTGCCCCCTG GAAGCCCTGG ATCTGAGCAC GGACATGACC
961 CAGCCACTGA GTGCCTACTT CATCAACTCC TCGCATRACA CCTATCTCAC TGCGGGGAGC
1021 CTGGCTGGGA CCTCGTGGT GGAGATGTAC CGCCAGGCAC TACTATGGGG CTGCCGCTGC
1081 GTGGAGCTGG ACGTGTGGAA GGGACGGCCG CCGTGGAGAG AACCTTCAT TACCCACGGC
1141 TTCACCATGA CCACAGAGGT GCCTCTGCGC GACGTGCTGG AGGCCATTGC CGAGACTGCG
65 1201 TTAAGACCT CGCCCTACCC CGTCACTCCT TCCTTCGAGA ACCATGTGGA CTCGGCAAAG
1261 CAACAGGCAA AGATGGCTGA GTACTGCCGC TCCATCTTTG GAGACGCGCT ACTCATCGAG

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1321 CCTCTGGACA AGTACCCGCT GGCCCCAGGC GTTCCCTGTC CCAGCCCCCA GGACCTGATG
1381 GGCCGTATCC TGGTGAAGAA CAAGAAGCGG CACCGACCCA GCGCAGGTGG CCCAGACAGC
1441 GCCGGGCGCA AGCGGCCCTT GGAGCAGAGC AATTCTGCCC TGAGCGAGAG CTCCGCGGCC
1501 ACCGAGCCCT CCTCCCCGCA GCTGGGGTCT CCCAGCTCTG ACAGCTGCCC AGGCCTGAGC
1561 AATGGGGAGG AGGTAGGGCT TGAGAAGCCC AGCCTGGAGC CTCAGAAGTC TCTGGGTGAC
1621 GAGGGCTGA ACCGAGGCC CTATGTTCTT GGACCTGCTG ACCGTGAGGA TGAGGAGGAA
1681 GATGAGGAAG AGGAGGAACA GACAGACCCC AAAAAGCCAA CTACAGATGA GGGCACAGCC
1741 AGCAGCGAGG TGAATGCCAC TGAGGAGATG TCCACGCTTG TCAACTACAT CGAACCTGTC
1801 AAGTTCAAGT CCTTTGAGGC TGCTCGAAAG AGGAACAAAT GCTTCGAGAT GTCGTCCTTT
1861 GTGGAGACCA AGGCCATGGA GCAACTGACC AAGAGCCCCA TGGAGTTTGT GGAATACAAC
1921 AAGCAGCAGC TCAGCCGCAT CTACCCCAG GGCACCCGCG TGGACTCCTC CAACTACATG
1981 CCCCAGCTCT TCTGGAACGT AGGGTGCCAG CTGTGTCGCG TCAACTTCCA GACCCTCGAT
2041 GTGGCGATGC AGCTCAACGC GGGCGTTTTT GAGTACAACG GCGCGAGCGG GTACCTGCTC
2101 AAGCCGAGT TCATGCGGGG GCCGGACAAG TCCTTCGACC CCTTCACTGA GGTATCGTG
2161 GATGGCATCG TGGCCAATGC CTTGCGGGTC AAGGTGATCT CAGGGCAGTT CCGTCCGAC
2221 AGGAAGGTGG GCATCTACCT GGAGGTGGAC ATGTTTGGCC TCCCTGTGTA TACGCGCGC
2281 AAGTACCGCA CCCGACCTC TCAGGGGAAC TCGTTCAACC CCGTGTGGGA CGAAGAGCCC
2341 TTCGACTTCC CCAAGGTGGT GCTGCCCACG CTGGCTTCAC TTCGATTGC AGCCTTTGAG
2401 GAGGGGGGTA AATTGCTAGG GCACCGGATC CTGCCTGTCT CTGCCATCCG CTCCGGATAC
2461 CACTACGTCT GCCTGCGGAA CGAGGCCAAC CAACCGCTGT GCCTGCCGGC CCGTCTCATC
2521 TACACCGAAG CCTCGGACTA CATTCTGTAC GACCACCAGG ACTATGCGGA GCGAGCTGGC
2581 AACCCCATTA AGCAGCTCAG CCTGATGGAC CAGAGGGCCC GGCAGCTGGC CGCCCTCATT
2641 GGGGAGAGTG AGGCTCAGGC TGGCCAAGAG ACGTGCCAGG ACACCCAGTC TCAGCAGCTG
2701 GGGTCTCAGC CGTCTCAA CCACCCCCC AGCCCACTGG ATGCCTCCCC CCGCCGGCCC
2761 CCTGGCCCCA CCACCTCCCC TGCCAGCACC TCCCTCAGCA GCCCAGGGCA GCGTGATGAT
2821 CTCATCGCCA GCATCCTCTC AGAGGTGGCC CCCACCCCGC TGGATGAGCT CCGAGGTAC
2881 AAGGCTCTGG TCAAGCTCCG GAGCCGGCAA GAGCGAGACC TGGGGAGCT GCGCAAGAAG
2941 CATCAGCGGA AGGCAGTCAC CCTCACCCGC CGCTGCTGG ATGGCCTGGC TCAGGCACAG
3001 GCTGAGGGCA GGTGCGGGT GCGGCCAGGT GCCCTAGGTG GGGCCGCTGA TGTGGAGGAC
3061 ACGAAGGAGG GGGAGGACGA GGCAAGCGG TATCAGGAGT TCCAGAACAG ACAGGTGCAG
3121 AGCCTGCTGG AGCTGCGGGA GGCCAGGTG GACGCAGAGG CCCAGCGGAG GCTGGAACAC
3181 CTGAGACAGG CTCTGCAGCG GCTCAGGGAG GTGCTCCTTG ATGCAAACAC AACTCAGTTC
3241 AAGAGGCTGA AAGAGATGAA CGAGAGGGAG AAGAAGGAGC TGCAGAAGAT CCTGGACAGA
3301 AAGCGCCATA ACAGCATCTC GGAGGCCAAG ATGAGGGACA AGCATAAGAA GGAGGCGGAA
3361 CTGACGGAGA TTAACCGTGC GCACATCACT GAGTCAGTCA ACTCCATCCG TCGGCTGGAG
3421 GAGGCCCAGA AGCAGCGGCA TGACCGTCTT GTGGCTGGGC AGCAGCAGGT CCGTCAACAG
3481 CTGGCAGGAG AGGAGCCCAA GCTGCTGGCC CAGCTGGCCC AGGAGTGTCA GGAGCAGCGG
3541 GCGAGGCTCC CCCAGGAGAT CCGCGGAGC CTGCTGGGCG AGATGCCGGA GGGGCTGGGG
3601 GACGGGCTC TGGTGGCCTG TGCCAGCAAC GGTACGACAC CCGGAGCAG CGGGCACCTG
3661 TCGGGCGCTG ACTCGGAGAG CCAGGAGGAG AACACGCAGC TCTGA

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GENBANK ID: D13119.1

DEFINITION HOMO SAPIENS P2 MRNA FOR ATP SYNTHASE SUBUNIT C, COMPLETE CDS.

VERSION D13119.1 GI:285909

CDS 31..456

/CODON_START=1

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1 TCTCTGCCA CAGCTCTCA CCCCCTGAAA ATGTTGCGCT GCTCCAAGTT TGTCTCCACT
61 CCCTCCTTGG TCAAGAGCAC CTCACAGCTG CTGAGCCGTC CGCTATCTGC AGTGGTGCTG
121 AAACGACCGG AGATACTGAC AGATGAGAGC CTCAGCAGCT TGGCAGTCTC ATGTCCCTTT
181 ACCTCACTTG TCTCTAGCCG CAGCTTCCAA ACCAGCGCCA TTTCAGGGA CATCGACACA
241 GCAGCCAAAGT TCATTGGAGC TGGGGCTGCC ACAGTTGGGG TGGCTGGTTC TGGGGCTGGG
301 ATTGGAAGT TGTTTGGGAG CCTCATCAT GGTATGCCA GGAACCTTC TCTGAAGCAA
361 CAGCTCTTCT CCTACGCCAT TCTGGGCTTT GCCCTCTCGG AGGCCATGGG GCTCTTTTGT
421 CTGATGGTAG CTTTCTCAT CCTCTTTGCC ATGTGAAGGA GCCGTCTCCA CCTCCATAG
481 TTCTCCGCG TCTGGTTGGC CCCGTGTGTT CCTTTTCTTA TACCTCCCCA GGCAGCCTGG
541 GGAACGTGGT TGGCTCAGGG TTTGACAGAG AAAAGACAAA TAAATACTGT ATTAATAAG

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GENBANK ID: NM_004530.1

DEFINITION HOMO SAPIENS MATRIX METALLOPROTEINASE 2 (GELATINASE A, 72KD GELATINASE, 72KD TYPE IV COLLAGENASE) (MMP2), MRNA.

VERSION NM_004530.1 GI:11342665

CDS 290..2272

/CODON_START=1

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 121 CGGGGGCGGG ACCATGAGCC GCTGAGCCGG GCAAACCCCA GGCCACCGAG CCAGCGGACC
 181 CTCGGAGCGC AGCCCTGCGC CGCGGACCAG GCTCCAACCA GCGGCGGAGG CGGCCACAGC
 241 CACCGAGCCA GCGACCCCGG GGGGACGCGC GGGGCCAGGG AGCGCTACGA TGGAGGCGCT
 301 AATGGCCCGG GCGCGCTCA CGGTCCCCT GAGGGCGCTC TGTCTCTGG GCTGCTGCT
 361 GAGCCACGCC GCCGCGCGCG CGTCGCCAT CATCAAGTTC CCCGCGATG TCGCCCCCAA
 421 AACGGACAAA GAGTTGGCAG TGCAATACCT GAACACCTTC TATGGCTGCC CCAAGGAGAG
 481 CTGCAACCTG TTTGTGCTGA AGGACACACT AAAGAAGATG CAGAAGTTCT TTGGACTGCC
 541 CCAGACAGGT GATCTTGACC AGAATACCAT CGAGACCATG CGGAAGCCAC GCTGCGGCAA
 601 CCCAGATGTG GCCAACTACA ACTTCTTCCC TCGCAAGCCC AAGTGGGACA AGAACCAGAT
 661 CACATACAGG ATCATTGGCT ACACACCTGA TCTGGACCCA GAGACAGTGG ATGATGCCCT
 721 TGCTCGTGCC TTCCAAGTCT GGAGCGATGT GACCCCACTG CGGTTTTCTC GAATCCATGA
 781 TGGAGAGGCA GACATCATGA TCAACTTTGG CCGCTGGGAG CATGGCGATG GATACCCCTT
 841 TGACGGTAAG GACGGACTCC TGGCTCATGC CTTCGCCCCA GGCACCTGGT TTGGGGGAGA
 901 CTCCCATTTT GATGACGATG AGCTATGGAC CTTGGGAGAA GGCCAAGTGG TCCGTGTGAA
 961 GTATGGCAAC GCCGATGGGG AGTACTGCAA GTTCCCCTTC TTGTCAATG GCAAGGAGTA
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 1441 CCGCAAGTGG GGCTTCTGCC TGGAGCACTC CCAAGACCCT GGGGCCCTGA TGGCACCCTA
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 1561 TTACACCTAC ACCAAGAACT TCCGTCTGTC CACCGGCCCC ACCCCACAC TGGGCCCTGT
 1621 CTATGGGGCC TCTCTGACA TTGACCTTGG ATTTGATGGC ATCGCTCAGA TCCGTGGTGA
 1681 CACTCCTGAG ATCTGCAAA AGGACATTGT ATTTGATGGC ACGCCACGTG ACAAGCCCAT
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 1921 AGCCAGCACC CTGGAGCGAG GGTACCCCAA GCCACTGACC AGCCTGGGAC TGCCCCCTGA
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 2101 GCTCATCGCA GATGCCTGGA ATGCCATCCC CGATAACCTG GATGCCGTG TGGACCTGCA
 2161 GGGCGGCGGT CACAGCTACT TCTTCAAGGG TGCCATTATG CTGAAGCTGG AGAACCAAG
 2221 TCTGAAGAGC GTGAAGTTTG GAAGCATCAA ATCCGACTGG CTAGGCTGCT GAGCTGGCCC
 2281 TGGCTCCAC AGGCCCTTCC TCTCCACTGC CTTGATACA CCGGGCCTGG AGAATAGAG
 2341 AAGGACCCCG AGGGGCTGG CAGCCGTGCC TTCAGTCTA CAGCTAATCA GCATTCTCAC
 2401 TCCTACCTGG TAATTTAAGA TTCCAGAGAG TGGCTCCTCC CGGTGCCCAA GAATAGATGC
 2461 TGACTGTACT CCTCCCAGGC GCCCTTCCC CCTCCAATCC CACCAACCCT CAGAGCCACC
 2521 CCTAAAGAGA TCCTTTGATA TTTTCAACGC AGCCCTGCTT TGGGCTGCC TGGTGTGCC
 2581 ACACCTCAGG CTCTTCTCCT TTCACAACCT TCTGTGGCTC ACAGAACCTT TGGAGCCAAT
 2641 GGAGACTGTC TCAAGAGGGC ACTGGTGGCC CGACAGCTG GCACAGGGCA GTGGGACAGG
 2701 GCATGGCCAG GTGGCCACTC CAGACCCCTG GCTTTTCACT GCTGGCTGCC TTAGAACCTT
 2761 TCTTACATTA GCAGTTTGCT TTGTATGCAC TTTGTTTTTT TCTTTGGGTC TTGTTTTTTT
 2821 TTTCCACTTA GAAATTGCAT TTCCTGACAG AAGGACTCAG GTTGTCTGAA GTCACTGCAC
 2881 AGTGATCTCT AGCCACATA GTGATGGTTC CCTGTTCAC TCTACTAGC ATGTCCCTAC
 2941 CGAGTCTCTT CTCCACTGGA TGGAGGAAAA CCAAGCCGTG GCTTCCCGCT CAGCCCTCCC
 3001 TGCCCTCTCC TTCAACCATT CCCCATGGGA AATGTCAACA AGTATGAATA AAGACACCTA
 3061 CTGAGTGGC

GENBANK ID: NM_000852.2
 DEFINITION HOMO SAPIENS GLUTATHIONE S-TRANSFERASE PI (GSTP1), MRNA.
 VERSION NM_000852.2 GI:6552334
 CDS 30..662

1 GGAGTTTCGC CGCCGACGTC TTCGCCACCA TGCCGCCCCA CACCGTGGTC TATTTCCAG
 61 TTCGAGGCCG CTGCGCGGCC CTGCGCATGC TGCTGGCAGA TCAGGGCCAG AGCTGGAAGG
 121 AGGAGGTGGT GACCGTGGAG ACGTGGCAGG AGGGCTCACT CAAAGCCTCC TGCCTATACG
 181 GGCAGTCCCC CAAGTTCCAG GACGGAGACC TCACCTGTGA CCAGTCCAAT ACCATCTGCG
 241 GTCACCTGGG CCGCACCCCT GGGCTCTATG GGAAGGACCA GCAGGAGGCA GCCCTGGTGG
 301 ACATGGTGAA TGACGGCGTG GAGGACCTCC GCTGCAATA CATCTCCCTC ATCTACACCA
 361 ACTATGAGGC GGGCAAGGAT GACTATGTGA AGGCACTGCC CCGGCAACTG AAGCCTTTTG
 421 AGACCTTGCT GTCCAGAAC CAGGGAGGCA AGACCTTCAT TGTGGGAGAC CAGATCTCCT

481 TCGCTGACTA CAACCTGCTG GACTTGCTGC TGATCCATGA GGTCCCTAGCC CCTGGCTGCC
541 TGGATGCGTT CCCCCTGCTC TCAGCATATG TGGGGCGCCT CAGCGCCCGG CCCAAGCTCA
601 AGGCCTTTCTT GGCCTCCCTT GAGTACGTGA ACCTCCCAT CAATGGCAAC GGGAAACAGT
661 GAGGGTTGGG GGGACTCTGA GCGGGAGGCA GAGTTTGCCT TCCTTTCTCC AGGACCAATA
721 AAATTTCTAA GAGAGCT

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GENBANK ID: XM_016524.4
DEFINITION HOMO SAPIENS CREATINE KINASE, MITOCHONDRIAL 1 (UBIQUITOUS)
(CKMT1), MRNA.
10 VERSION XM_016524.4 GI:17477504
CDS 358..1704
/CODON_START=1

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1 CGCGCGAGTC TCAGGTCCCC CTAATTACCT GCGGGGTGCT GCCCACCCTT GCCTTCGCGC
61 ACCTAGCGCG GTGGCAGGCG GGAAGGCGGG GCCTGGGGGA GCCCACCCTT TGGAGACTGC
121 GGCTGGGGCC TCCCTCTCCT CCGCCCCCCC GCCTGCCACT AGCTCATTGC GCCTCTCTCT
181 CAGTCTGATT GGACCCGGCT CCCATTCCGG CTCCAGCCTC CAATCCGACC CCCATTTCGG
241 CTGCAGCCTC GGACCTAGCT CCGGCCCTCG GTCTATCCGG TTGCATCCTC CCTCCCTGTT
301 CCGGATCTTA TCTTGCGCCA GCGCCTACTC CAGGATCCCG TAGCCAGACC TCAAGCCATG
361 GCTGGTCCCT TCTCCCGTCT GCTGTCCGCC CGCCCCGGAC TCAGGCTCCT GGCTTTGGCC
421 GGAGCGGGGT CTCTAGCCCG TGGGTTTCTG CTCGACCCGG AACCTGTACG AGCTGCCAGT
481 GAACGACGGA GGCTGTATCC CCCGAGCCAG ACATGGCCAA CTGGACAGCT CCCAGGTAAC
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601 GCTGAGTACC CAGACCTCCG AAAGCACAAC AACTGCATGG CCAGTCACCT GACCCCAGCA
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781 GATGAGGAGA CCTATGAGGT ATTTGCTGAC CTGTTTGACC CTGTGATCCA AGAGCGACAC
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961 CGAGGACTCA GTCTGCCTCC AGCTTGCACT CGAGCAGAGC GACGAGAGGT GGAACGTGTT
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1141 GTGTCCCGT TGCTGACTGC AGCAGGAATG GCTCGAGACT GGCCAGATGC TCGTGGAATT
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1501 CAAAAACGTG GTACTGGAGG AGTGGACACT GCTGCTACAG GCGGTGCTCT TGATATTTCT
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GENBANK ID: NM_001443.1
DEFINITION HOMO SAPIENS FATTY ACID BINDING PROTEIN 1, LIVER (FABP1), MRNA.
VERSION NM_001443.1 GI:4557576
CDS 43..426

1 AGAGCCGCAG GTCAGTCGTG AAGAGGGAGC TCTATTGCCA CCATGAGTTT CTCCGGCAAG
61 TACCAACTGC AGAGCCAGGA AAACTTTGAA GCCTTCATGA AGGCAATCGG TCTGCCGGAA
121 GAGCTCATCC AGAAGGGGAA GGATATCAAG GGGGTGTCGG AAATCGTGCA GAATGGGAAG
181 CACTTCAAGT TCACCATCAC CGCTGGGTCC AAAGTGATCC AAAACGAATT CACGGTGGGG
241 GAGGAATGTG AGCTGGAGAC AATGACAGGG GAGAAAGTCA AGACAGTGGT TCAGTTGGAA
301 GGTGACAATA AACTGGTGAC AACTTCAAA AACATCAAGT CTGTGACCGA ACTCAACGGC
361 GACATAATCA CCAATACCAT GACATTGGGT GACATTGTCT TCAAGAGAAT CAGCAAGAGA
421 ATTAAACAA GTCTGCATTT CATATTATTT TAGTGTGTAA AATTAATGTA ATAAAGTGAA
481 CTTTGTGTTT

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GENBANK ID: NM_001220.1
DEFINITION HOMO SAPIENS CALCIUM/CALMODULIN-DEPENDENT PROTEIN KINASE (CAM
KINASE) II BETA (CAMK2B), MRNA.
VERSION NM_001220.1 GI:10835005
CDS 47..1675

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1 GCGGCCCGCG TCGACCGAGC GCACGCCGAG CCCGTCCGCC GCCGCCATGG CCACCACGGT
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121 CTCTGTGGTC CGACGCTGTG TCAAGCTCTG CACCGGCCAT GAGTATGCAG CCAAGATCAT
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DEFINITION HOMO SAPIENS ATPASE, NA+/K+ TRANSPORTING, BETA 1 POLYPEPTIDE
(ATP1B1), MRNA.
VERSION NM_001677.1 GI:4502276
CDS 127..1038

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VERSION U82535.1 GI:2149155
CDS 36..1775
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VERSION NM_000789.1 GI:4503272
CDS 23..3943

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GENBANK ID: XM_052144.2

DEFINITION HOMO SAPIENS APOLIPOPROTEIN A-IV (APOA4), MRNA.

VERSION XM_052144.2 GI:15314431

CDS 114..1304

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GENBANK ID: M29366.1

DEFINITION HUMAN EPIDERMAL GROWTH FACTOR RECEPTOR (ERBB3) MRNA, COMPLETE CDS.

VERSION M29366.1 GI:181979

CDS 100..4128

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3961 GGGTATGAAG AGATGAGAGC TTTTCAGGGG CCTGGACATC AGGCCCCCA TGTCCATTAT
4021 GCCCGCCTAA AACTCTACG TAGCTTAGAG GCTACAGACT CTGCCTTGA TAACCTTGAT
4081 TACTGGCATA GCAGGCTTTT CCCCAAGGCT AATGCCCAGA GAACGTAAC CTGTCTCCCT
4141 GTGGCACTCA GGGAGCATT AATGGCAGCT AGTGCCCTTA GAGGGTACCG TCTTCTCCCT
15 4201 ATTCCCTCTC TCTCCAGGT CCCAGCCCTT TTTCCCACT CCCAGACAAT TCCATTCAAT
4261 CTTTGGAGGC TTTTAAACAT TTTGACACAA AATTCTTATG GTATGTAGCC AGCTGTGCAC
4321 TTTCTTCTCT TTCCCAACCC CAGGAAAGGT TTTCTTAT TTTGTGTGCT TCCCACTCCC
4381 ATTCTCAGC TTCTTCAGC GCACCTCTGG AGATATGAAG GATTACTCTC CATATCCCTT
4441 CCTCTCAGC TCTTGACTAC TTGGAAC TAGCTCTTATGT GTGCCTTTGT TTCCCATCAG
20 4501 ACTGTCAAGA AGAGGAAAGG GAGGAAACCT AGCAGAGGAA AGTGTAAATT TGGTTTATGA
4561 CTCTTAACCC CCTAGAAAGA CAGAAGCTTA AAATCTGTGA AGAAGAGGT TAGGAGTAGA
4621 TATTGATTAC TATCATAAT CAGCACTTAA CTATGAGCCA GGCATCATAC TAAACTTCAC
4681 CTACATTATC TCACCTTAGT CTTTATCATC CTTAAACAA TTCTGTGACA TACATATTAT
4741 CTCATTTTAC ACAAAGGGA GTCGGCATG GTGGCTCATG CCTGTAATCT CAGCACTTTG
25 4801 GGAGGCTGAG GCAGAGGAT TACCTGAGC AAGAGTTT AGACCAGCT AGCCACATA
4861 GTAAGACCCC CATCTCTT

30 GENBANK ID: X54686
VERSION X54686.1 GI:56909

35 MCTKMEQAFYHDDSYAAAGYGRSPGSLSLHDYKLLKPTLALNLA
DPYRGLKPGARGPGPEGSGAGSYFSGQGS DTGASLKLASTELERLIVPNSNGVITTT
PTPPGQYFPRGGSGGTGGGVTEEQEGFADGFVKALDDLQKMNHVTPPNVSLGASG
GPOAGPGGVYAGPEPPPVYTNLSSYPASAPSGSGTAVGTGSSYPTATISYLPHAPP
FAGGHPAQLGLSRGASAFKEEPQTVPEARSRDATPPVSPINMEDQERIKVERKRLNR
LAATKCRKRLERLARLEDKVKTLKAENAGLSSAAGLLREQVAQLKQKVMTHVSNCGQ
LLLGVKGHAF

40 GENBANK ID: D26307
VERSION D26307.1 GI:450471

45 METPFYGEELSLAAGASSVAGAAGAPGGGGFAPPGRAFPAGP
PTSSMLKKDALTLSLAEQGAAGLKPGSATAPASALRPDGPDLGLLLKSLASPE
LERLIQSNGLVTTPTSTQFLYPKVAASEEQEFAEGFVKALDLHKQSLGAATAAT
SGAPAPPADLAATPGATETPVYANLSSFAGGAGPPGGAATVAFAAEPVFPFPPGA
LGPPPPPHPRLAALKDEPQTVPDVPSFGDSPPLSPIDMDTQERIKAEKRLNRNIAA
SKCRKRLERISRLKEKVKTLKSONTELASTASLLREQVAQLKQKVLSHVNSGCQLLP
QHQPAY

50 GENBANK ID: NM_012747.1
VERSION NM_012747.1 GI:6981591

55 MAQWNQLQQLDTRYLEQLHQLYSDSFMELRQFLAPWIESQDWA
YAASKESHATLVFHNLLGEIDQYSRFLQESNVLYQHNLRRIKQFLQSRYLEKPMIEIA
RIVARCLWEESRLQTAATAAQGGQANHTAAVTEKQMLEQHLQDVRKRVQDLEQ
KMKVVENLQDDFDENYKTLKSQGDMDLNGNNSVTRQKMQOLEQMLTALDQMRISIV
SELAGLLSAMEYVQKTLTDEELADWKRQIACIGGPPNICLDRLENWITSLAESQLO
TRQIKKLEELQKQVSYKGDPIVQHRPMLERIVDLFRNLKSAFVVERQPCMPMHPD
RPLVIKTGVQFTTKVRLLVKFPPELNYQLKIKVCI DKDSGDVAALRGSRKFNLGTNTK
60 VMNMEESNNGSLSAEFKHLTLREQRCNGGRANCASLIVTEELHLITFETEVYHQGL
KIDLETHSLPVVISNICOMPNAWASILWYNMLTNNPKNVNFTKPIGTWDQVAEVL
SWQFSSTTKRGLSIEQLTLAEKLLGPGVNYSGCQITWAKFCKENMAGKGFVFWWLD
NIIDLVKYILALWNEGYIMGFISKERERAILSTKPPGTFLLRFSESSKEGGVFTWV
EKDISGKTQIQSVEPYTKQQLNNMSFAEIMGYKIMDATNILVSLVLYLPDIPKEEA
65 FGKYCRPESQEHPEADPGSAAPYLKTKFCVTPPTCSNTIDLPMSPRTLDLSMQFGNN
GEGAEPESAGGQFESLTFMDLTSECATSPM

GENBANK

ID: L26267.1
VERSION L26267.1 GI:425471

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REILNPPEKETQGEPSLFMASTKTEAIAPASTMEDKEEDVGFO
DNLFLEKALQIAKRHANALFDYAVTGDVKMLLAVQRHLTAVQDENGDSVLHLAIHLH
AQLVRDLLEVTSGSISDDIINMRNDLYQTPLHLAVITKQEDVVEDLLRVGADLSLLDR
WGNSVLHLAAKEGHDKILGVLLKNSKAALLINHPNGEGLNAIHAVMSNSLSCLQLLV
AAGAEVNAQEQSGRTALHLAVEYDNISLAGCLLLEGDALVDSTTYDGTTPHIAAGR
GSTRLAALLKAAGADPLVENFEPLYDLDSDWEKAGEDEGVVPGTTPDMAANWQVFDI
LNGKPYEPVFTSDDILPQGDIKQLTEDTRLQLCKLLEIPDPKNWATLAQKLGILN
NAFRLSPAPSKTFLMDNYEVSGGTIKELVEALRQMGYTEAEVIAAFRTPETTASSPV
TTAQAHLLPLSSSTRQHIDELRDNSVCDSGVETSFRLKSFSESLTGDGPLLNLNM
PHNYGQDGPPIEGKI

GENBANK ID:

M34356.1
VERSION M34356.1 GI:181042

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MTMESGAENQQSGDAVTEAENQOMTVQAQFQIATLAQVSMPPA
HATSSAPTVTVLQVLPNGQTVQVHGVIQAQPSVIQSPQVQTVQISTIAESQESVD
SVTDSQKRREILSRPSYRKILNDLSSDAPGVPRIEEEKSEETSAPAITTTVPTPI
YQTSSGQYIAITQGGAIQLANNGTDGVQGLQTLTMTNAATQPGTTILQYAQTTDGOQ
ILVPSNQVVQAASGDVQTYQIRTAFTSTIAPGVVMASSPALPTQPAEEAARKREVRL
MKNREARECRKKKEYVKLENRVAVLENQNKTLIEELKALKDLYCHKSD

GENBANK ID:

X68193.1
VERSION X68193.1 GI:53353
MANLERTFIAIKPDGQVQGLVGEIIRFEQKGFRLVAMKFLRAS
EEHLKQHYIDLKDRPFFPGLVKYMNSGPVVAMVWEGNLVVKTRVMLGETNPADSKPG
TIRGDFCIQVGRNIIHGSDSVESAEKEIHLWFKPEELIDYKSCAHDWVYE

GENBANK

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ID: U29200.1
VERSION U29200.1 GI:924934

MANLERTFIAIKPDGQVQ

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GENBANK ID: L35572
VERSION L35572.1 GI:531219

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MMVHCAGCERPILDRFLNLVLDRAWHIKCVQCCECKTNLSEKCF
SREGKLYCKNDFRRFGTKCAGCAQGISPSDLVRKARSKVFHLNCFCTMCVNKQLSTG
EELYVIDENKFCKDDYLSSSLKEGSLNSVSSCTDRSLSPDLQDPLQDDPKETDNST
SSDKETANNENEEQNSGTRRGFRTTIKAKQLETLLKAAFAATPKPTRHIREQLAQETG
LNMVRVQVWFQNRRSKERRMKQLSALGARRHAFFRSRRMRPLGGRLESEMLGSTPY
TYYGDYQSDYYAPGGNYDFFAHGPPSQAQSPADSSFLAASGPGSTPLGALEPPLAGPH
GADNPRFTDMISHPDTPSPPEGLPGALHPMPGEVFSGGPSPFPMSGTSGYSGPLSHP
NPELNEAAVW

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GENBANK ID: X55787.1
VERSION X55787.1 GI:296022
MKAADVLDKPTLTIIKTEKVDLELFPSPMECADVPLSTPSSKEM

60

MSQALKSYSGFTKEQRRGIPKDPROWTDTHVRDWMWAVNEFSLKGVDFHKFCMSG
AAVCALGKECFLELAPDFVGDILWEHLEILQKEDVKPYQVNGANPTYPESCYTSVYFI
SYGIEHAQCVPPSEFSEPSFITESYQTLHPISSSELLSLKYENDYPSVILRDPLQET
LQTDYFRIKQEVLTFPNNMCLGRASRGKLGQDSFESVESYDSCDRLTQSWSSQSFNS
LQRVPSYDSFDYEDYPAALPNHKKPGTKFDYVRDRADLNKDKPVIAPAAAGYTGSGP
IQLWQFLLELLTDKSCQSFISWTGDCWEFKLSDPDEVARRWGKRKNKPSMNYEKLRSRA
LRYYYDKNIIHKTAGNAYVYAFVCDLQSLGTYPEELHAMLVDVKPDAD

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GENBANK ID: M11507.1
DEFINITION Human transferrin receptor

VERSION M11507.1 GI:339515

MMDQARSAFSNLFGGEPLSYTRFSLARQVDGDNSHVEMKLAVDE
EENADNNTKANVTTPKRCSGSICYGTIAVIVFFLIGFMIGYLGCKGVEPKTECERLA
GTESPVREEPEGEDFPAARRLYWDDLKRKLSEKLDSTDFSTIKLLNENSYPREAGSQ
KDNELALYVENQFREFKLSKVWRDQHFVKIQVKDSAQNSVIIVDKNGRLVYLVENPGG
YVAYSKAATVTGKLVHANFGTKKDFEDLYTPVNGSIVIVRAGKITFAEKVANAESLNA
IGVLIYMDQTKFPIVNAELSFFGHAHLGTGDPYTPGFPSFNHTQFPPSRSSGLENIPIV
QTISRAAAEKLFGNMEGDCPSDWKTDSTCRMVTSSEKNVKTIVSNVLKEIKILNIFGV
IKGFVEPDHYVVVGAORDAWGPGAAGSGVGTALLLKLQMFSDMVLKDGFPQPSRSIIF
ASWSAGDFGSGATEWLEGYLLSSHLKAFYINLDKAVLGTSNFKVSASPLLYTLIEK
TMQNVKHPVTGQFLYQDSNWASKVEKLTLDNAAPFLAYSGIPAVSFCECDTDYPYL
GTTMDTYKELIERIPELNKVARAAAEVAGQFVIKLTHDVELNLDYERYNSQLLSFVRD
LNQYRADIKEMGLSLQWLYSARGDFFRATSLTTDFGNAEKTDRFVMKLNDRVMRVE
YHFLSPYVSPKESPFPHVFWGSGSHTLPALLENLKLKRNQNGAFNETLFRNQLALATW
TIQGAANALSGDVWDIDNEF

GENBANK ID: AAA60255

VERSION AAA60255.1 GI:190927

1 mteyklvvvg agvgvgsalt iqliqnhfvd eydptiedsy rkqvvidget clldildtag
61 qeysamrdq ymrtgegflic vfainnsksf adinlyrequ krkdsddvp mvlvgnkcdl
121 prtvtvkqa helaksyqip fietsaktrq gvedafytlv reirqyrmkk lnsddgtqg
181 cmglpcvmm

GENBANK ID: AAB38309

VERSION AAB38309.1 GI:1497931

1 mslvndlli ccrqlehdra terkkevekf krlirdpeti khldrhsdsk qgkylndwav
61 frflqkyiqk etecclriak nvsastqasr qkkmqeissl vkyfikcanr raprlkccgel
121 lnyimdtvkd ssngaiygad csnillkdil svrkywceis qqqwlelfsv yfrlylkpsq
181 dvhrvlvari ihavtkgccc qtdglnskfl dffskaiqca rgeksssgln hilaaltifl
241 ktlavnfriar vcelgdeilp tlliywtqhr lndslkevii elfqlqiyih hpkqaktqek
301 gayestkwsr ilynydillv neishigsg kyssgfrnia vkenlielma dichqvfned
361 trsleisqsy tttqressdy svpckrkkie lgwevikdhl qksqndfdlv pwlqiatqli
421 skypaslpnc elspilmils qlipqqrhge rtpyvrlcrl evalcqdkrs nlessqksdl
481 klwnkiwci tfrgisseqi qaeafgllga iaggslvevd refwklftgs acrpsscavc
541 cltlatlttsi vpgtvkmgie qnmcevnrsf slkesimkw lfyqllegdle nstevppilh
601 snfphlvlek ilvsltmknc kaamnnffqsv pecehhgkdk eelsfsevee lflqttfdkm
661 dfltivrecg lekhsqsigf svhgnlkesl drcllglseq llnnyssseit nsetlvrcsr
721 llvgvlvcyc ymgviaeaea ykselfgkak gfflrlttsk lmnadiadick slasfikkpf drgevesmed
781 qlctrcslnc tkkspnkias gfflrlttsk lmnadiadick slasfikkpf drgevesmed
841 dtngnlmeve dqssmnlfn ypdssvsdan epgesqstig ainplaeeyl skqdlflfmd
901 lkflclcvtt aqntvvsfra adirrkllml idsstleptk slhlmnylml lkelppgeey
961 lpmadvlell kplsncvcsly rrdqdvckti lnhvlhvkn lgqsnmdsen trdaqgqfl
1021 vigafwhltk erkyifsvrm alvnciktll eadpyskwai lnmvgkdfvp nevftqflad
1081 nhqvmrlaa esinrlfqdt kgdssrllka lplklqqtat enaylkageg mremshsaen
1141 petldeiynr ksvlltliav vlscspicek qalfalcksv kenglephlv kkvlekvset
1201 fgrrledfm ashldylvle wlnlqdteln lssfpfilln ytniedfyr cykvliplhv
1261 irshfdevks ianqigedwk slttdcfpki lvnilyfay egtrdsqmaq qretatkvyd
1321 mlksenllgk qidhlfisnl peivvellmt lhpanssas qstdlcdfsg dldpappnph
1381 fpshvikatf ayisnchktk lksileilsk spdsyqkill aiceqaetn nvykhrilk
1441 iyhlvsl11 kdiksglga wafvlrdviy tlihyinqrp scimdvslrs fslccdl1sq
1501 vcqtavtyck dalenhlhvi vgtliplvye qvevqkvld llkylvidnk dnenlyitik
1561 lldpfpdhvv fkdrlritqk ikysrgpfsl leehnlflsv svydalpltr leglkdlrrq
1621 lelhkdmvd imrasqdnpg dgimvklvn llqlskmain htgekevlea vgsclgevgp
1681 idfstiaiqh skdasyskal klfedkelqw tfimltylnn tlvedcvkvr saavtclkn
1741 latktghsfw eiymktdpm laylqpfrts rkkflevprf dkenpfegld dinlwiplse
1801 nhdiwiktlt cafldsggk ceilqlkpm cevkttdfcqt vlpylidhll lqdtneswrn
1861 llsthvqgff tscrlrhfsqt srsttpandl sesehffrci yadkksmddq ekrslafeeg
1921 rpsstg1fnd afwldllyle vakvaqscas hftallyaei yadkksmddq ekrslafeeg
1981 sqsttissls ekskeetgis lqdl1leiy sigepdslyg cgggkmlqpi trlrtyeha
2041 mwgkalvtyd letaipsstr qagiiqalqn lglchilsvy lkgldyenkd wcpel1elhy
2101 qaawrnmgwd hctsvskeve gtsyheslyn alqslrdref stfyelskya rvkeveemck
2161 rslesvysly ptlsrlqag elsesigelfs rsvthrlse vyikwqkhsq llkdsd1fsq
2221 epimalrtvi leilmekemd nsqrecikdi ltkhlvelsi lartfkntql peraifqikq

5 2281 ynsvscgvse wgleeaqvw akkeqslals ilkqmikkld ascaannpsl kityteclrv
 2341 cgnwlaetcl enpavimqty lekavevagn ydgessdelr ngkmkafisl arfsdtqyqr
 2401 ienymkssef enkqallkra keevgllreh kiqtntyrvk vgreleidel alralkedr
 2461 rflckaveny incllsgeeh dmwvfrlcs lwnsgvsev ngmmkrdgmk iptykflplm
 2521 yqlaarmgtk mmgglgfhev innlisrism dhphntlfii lalanandrde fltkpevarr
 2581 sritknvpkq ssqldedrte aanriictir srpgmvrsv ealcdayiil anldatgwkt
 2641 qrkginipad qpitklknl dvvvtmeik vdhtgeygnl vtiqsfkaef rlaggvnlpk
 2701 iidcvgsdgk errqlvkgrd dlrqdamvqq vfgmctllq rntetrkrkl tictykvvpl
 2761 sqrgsvlewc tgtvpigefl vnnedgahkr yrpndfsafq cqkkmmevqk ksfeekyevf
 10 2821 mdvcqnfqpv fryfcmekfl dpaiwfekrl aytrsvatss ivgyilglgd rhvqniline
 2881 qsaelvhlidl gvafeqqkil ptpetvpfrl trdivdgmgi tgvegfvrrc cektmevmrn
 2941 sqetlltive vliydlplfdw tmnpkaly l qqrpedetel hptlnaddqe ckrnlstdidq
 3001 sfdkvaervl mrlqekikgv eegtvlsvgg qvnlliqgai dpknlsrlfp gwkwaw
 //

20 GENBANK ID: AAA59145.1
 VERSION AAA59145.1 GI:307058
 1 mlkpslpfts llflqlpllg vglnttiltp ngnedttadf flttmptdsl svstlplpev
 61 qcfvfnvey m nctwnsssep qptnltlhyw yknsdndkvq kcshylfsee itsgcqqlqk
 121 eihlygtfiv qlqdprrpr qatqmlklqn lvipwapenl tlhlksesql elnwnnrfln
 181 hclehlvqyr tdwdhswteq svdyrhkfs l psvdgqkryt frvrsrfnpl cgsaqhwsew
 241 shpihwsnt skenpflfal eavvisvgsm gliisllcvy fwlermpri ptlknledlv
 301 teyhgnfsaw sgvsqklaes lqpdysrlc lvseippkkg algegpgasp cnqhsqpywap
 361 pcytlkpet

30 GENBANK ID: AAC50825.1
 VERSION AAC50825.1 GI:1117984
 1 mvaprplrrv vlfyqgklcs magnfwqssh ylwildkqd llkerqkdik flseeeywkl
 61 giffntviga lgehlklrqq viatatvyfk rfyarysiks idpvlmptc vflaskveef
 121 gvsntria aatsvlktrf syafpkfpy rmnhilecef yllelmdccl ivyhyprpl
 181 qyvqdmqged mlplawriv ndtyrtdlcl lyppfmiala clhvaccvqg kdarqwaef
 241 svdmekilei irvilklieq wknfderkem atilskmpkp kpppnsegeq gpngsqnssy
 301 sqs

40 GENBANK ID: AAC50473.1
 VERSION AAC50473.1 GI:1314346
 1 mamssggsgg gvpegedsvl frgtgqsd sdiwddtali kaydkavasf khalkngdic
 61 etsgkpkttp krkpakknks qkntaaslq qwkvgdkcsa iwsedgciyp atiasidfkr
 121 etcvvvytgy gnreeqnlsd llspicevan nieqnaqene nesqvstdes ensrspgnks
 181 dnkpkpsapw nsflpppppm pgprlpgkp glkfngpppp pppppphlls cwlpfpfsgp
 50 241 piippppic pdsldadal gsmliswysm gyhtgyymgf rqnqkegrcs hsln

55 GENBANK ID: CAC15525
 VERSION CAC15525.1 GI:11137517
 1 mvsrdqahlg pkyvglwdfk srtdeelsfr agdvfhvark eeqwwatll deaggavaqg
 61 yvphnylaer etvesepwff gcisrseavr rlqaegnats aflirvsekp sadyvlsvrd
 121 tqavrhykiw rraggrlhlh eavsfilspe lvnyhraqls shglrlaapc rkhepeplph
 181 wddwerpree ftlcrklgsg yfgevfeqlw kdrvqvaiqv isrdnllhqq mlqseiqlmk
 241 klrkhilal yavsvgdv yitelmakg sllellrdsd ekvlpvsell diawqvaegm
 301 cylesqnyih rdlaarnilv gentlckvgd fglarliked vylshdhnip ykwtapeals
 361 rghystksdv wsfqillhem fsrgqvpypp msnheafllrv dagyrmpcpl ecppswhkml
 421 ltcwcrdpeq rpcfkaler lssftsypen t

65

GENBANK ID: AAB84296
VERSION AAB84296.1 GI:2613135

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1 tstttvrglna strylfrvra svqglgdwsn tveettlglq saspvqesrv aedgldqqlv
61 lavvgsvsat cltilaalla lvcirrsclh rrhtftyqsg sgeetilqfs sgtltltrrp
121 kpqpeplsyp vle

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GENBANK ID: X63594.1
VERSION X63594.1 GI:57673
MFQAPAGHGQDWAMEGPRDGLKKERLVDDRHDSSGLDSMKDEDEYEQ
MVKELREIRLQPEAPLAEPWKQQLTEDGDSFLHLAIHEEKTILMEVIGQVKGDLA
FLNFQNNLQQTPLHLAVITNQPGIAEALLKAGCDELRDFRGNTPLHLACEQGCLASV
AVLTQTCTPQHLHSLVQATNYNGHTCLHLASINGYLGIVEHLVTLGADVNAQEPNCGR
TALHLAVDLQNPDLVSLLLKCGADVNRVTYQGYSPYQLTWGRPSTRIQQQLGQLTLEN
LQTLPESEDEESYDTESEFTEDELPHYDDCVFGGQRLTL

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GENBANK ID: AAB60641
VERSION AAB60641.1 GI:516515

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1 meqgdqsmke gmgttwillst pqhwmqqfy netyygrtge fmedfpltl1 wsvtvsmfpp
61 ggfigsllvg plvnkfgkrg allfnnifsi vpailmgcsr vatsfeliii srllvgicag
121 vssnvvpmyl gelapknlg algvvpqlfi tvgilvaqif glrnl1anvd gwpillgltg
181 vgaalqllll pffpespryl liqkkdeaaa kkalqtlrgw dsvdrevaei rqedeaekaa
241 gfisvklfr mrs1rwqls iivlmggqql sgvnaiyya dqiylsagvp eehvqyvtg
301 tgavnnvmtf cavfvvellg rrlllllgfs icliaccvlt aalalqdtvs wmpyisivcv
361 isyvighalg pspipallit eiflqssrps afmvvggsvhw lsnftvglif pfieglgpy
421 sfivfavicl lttiyifliv petkaktfie inqiftkmnk vsevypekee lkelpptse
481 q

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GENBANK ID: M29069
VERSION M29069.1 GI:205553

5 MLSCCTTSTMPGMICKNSDLEFDSLKPCFYPEDDDIYFGGRNSTP
PGEDIWKKFELLPTPRLSPGRALAEDSLEPANWATEMLLPEADLWSNPAAEEDI FGLK
GLSGSSSNPVVLQDCMWSGFSSREKPEVTVSEKLPGGCGSLAVGAGTLVPGAAAATSA
GHARSGTAGVGRRKAAWLTSLHLDSECVDSAVIFPANKRESMPVATIPASAGAAISL
10 GDHQGLSSSLEDFLSNSGYVEEGGEEIYVVMLGETQFSKTVTKLPTAAHSENAALTPE
CAQSGELILKRSDLIQEQHNYAAPPLPYAEDARPLKKPRSQDPLGPKCVLRPKAPRL
RSRSNSDLEDIERRRNHNRMERQRRDIMRSSFLNLRDLVPELVHNEKAAKVILKKAT
EYIHTLQTDSEKLLVEREKLYERKQQLLEKIKQSAVC

15 GENBANK ID: M29039.1
VERSION M29039.1 GI:186626

MCTKMEQPFYHDDSYTATGYGRAPGGLSLHDYKLLKPSLAVNLA
DPYRSLKAPGARGPGPEGGGGSYFSGQSDTGASLKLASSELERLIVPNSNGVITTT
PTPPGQYFYPRGGSGGGAGGAGGGVTEEOEGFADGFVKALDDLHKMNHVTPPNVSLG
20 ATGGPPAGPGGVYAGPEPPPVYTNLSSYS PASASSGAGAAVGTGSSYPTTTISYLPH
APPFAGGHPAQLGLGRGASTFKEEPQTVPEARSRDATPPVSPINMEDQERIKVERKRL
RNRLAATKCRKRRLERIRARLEDKVKTILKAENAGLSSTAGLLREQVAQLKQKVMTHVSN
GCQLLLGVKGHAF

25 GENBANK ID: X56681.1
VERSION X56681.1 GI:34018

30 METPFYGDALSGLGGGASGSGTFA SPGRLFPGAPPTAAAGSM
MKKDALTLSLSEQVAAALKPAPAPASYPPAADGAPSAAPPDGLLASPDLLKLASPE
LERLI IQSNGLVTTTPTSSQFLYPKVAASEEQEFAEGFVKALDDLHKMNHVTPPNVSLG
AAAAAAGGPGSGTATGSAPPGELAPAAAPEAPVYANLSSYAGGAGGAGGAATVAFAAE
35 PVFPFPPPPGALGPRLAALKDEPQTVPDVPSFGESPPLSPIDMTQERIKAEKRL
RNRIAASKCRKRRLERISRLLEKVKTLKSQNTLASTASLLREQVAQLKQKVLSHVNS
GCQLLPQHQPAY

40 GENBANK ID: NM_003150.1
VERSION NM_003150.1 GI:4507252

MAQWNQLQLDTRYLEQLHQLYSDSFPMELRQFLAPWIESQDWA
YAASKESHATLVFHNLLGEIDQQYSRFLQESNVLYQHNLRRIKQFLQSRYLEKPMIEA
RIVARCLWEESRLLQTAATAAQGGQANHPTAAVVTEKQQLMEQLQDVRKRVQDLEQ
45 KMKVVENLQDDDFDNFKTLKSQGMQDLNGNNSVTRQKMQQLEQMLTALDQMRISIV
SELAGLLSAMEYVQKTLTDEELADWKRRQIACIGGPPNICLDRLNWTSLAESQLQ
TRQQIKKLEELHQVSYKGDPIVQHREMLBERIVELFRNLKSAFVVERQPCMPMHPD
RPLVIKTGVQFTTKVRLLVKFPPELNYQLKIKVCIDKDSGDVAALRGSRKENILGTNTK
50 VMNMEESNNGSLSAEFKHLTLREQRCGNGGRANCDASLIVTEELHLITFETEVYHQL
KIDLETHSLSVVISNICQMPNAWASILWYNMLTNNPKNVNFTKPPIGTWDQVAEVL
SWQFSSTTKRGLSIEQLTLAEKLLGPGVNYSGCQITWANFCENMAGKGFYSWVWLD
NIIDLVKKYILALWNEGYIMGFISKEERAILSTKPPGTFLRRFSESSKEGGVFTTWV
EKDISGKTQIQSVEPYTKQQLNNMSFAEIIIMGYKIMDATNILLSPLVLYLPDIPKEEA
55 FGKYCRPESQEHPEADPGSAAPYLKTKFICVTPPTCSNTIDLPMSPRALDSLMOFGNN
GEGAEPGAGGQFESLTFDMELTSECATSPM

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(54) Title: GENETIC ANALYSIS OF PEYER'S PATCHES AND M CELLS AND METHODS AND COMPOSITIONS TARGETING PEYER'S PATCHES AND M CELL RECEPTORS

(57) Abstract: Methods of increasing or decreasing the levels of a protein in a PP cell; methods of increasing antigen, vaccine, DNA vaccine delivery to M cells, use of human serum albumin and other transport enhancing proteins to enhance oral drug delivery; use of calreticulin to enhance oral antigen delivery, use of other cell surface proteins, receptors, and transporters to enhance delivery to M cells of antigens or vaccine delivery vehicles, use of other cytoplasmic proteins to regulate intracellular trafficking and delivery to mucosal immune sampling and processing systems.

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INTERNATIONAL SEARCH REPORT

International Application No

PCT/IB 02/03866

A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 C12N15/12 C12N15/63 C07K14/00 A61K48/00 A61K47/42
A61K39/00 A61K47/48

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the International search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, MEDLINE, EMBASE, BIOSIS

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 6 060 082 A (CHEN HONGMING ET AL) 9 May 2000 (2000-05-09) column 1, line 11 - line 26 column 2, line 38 - line 47 column 2, line 66 - column 3, line 44 column 7, line 4 - line 33 column 19, line 31 - column 22, line 35 ----- -/--	1-58, 61-64



Further documents are listed in the continuation of box C.



Patent family members are listed in annex.

* Special categories of cited documents :

"A" document defining the general state of the art which is not considered to be of particular relevance

"E" earlier document but published on or after the international filing date

"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.

"&" document member of the same patent family

Date of the actual completion of the international search

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INTERNATIONAL SEARCH REPORT

International Application No

PCT/IB 02/03866

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	GULLBERG ELISABET ET AL: "Expression of specific markers and particle transport in a new human intestinal M-cell model." BIOCHEMICAL AND BIOPHYSICAL RESEARCH COMMUNICATIONS, vol. 279, no. 3, 29 December 2000 (2000-12-29), pages 808-813, XP002252802 ISSN: 0006-291X page 808 abstract page 810, right-hand column, paragraph 1 -----	59,60
A	US 6 117 632 A (O'MAHONY DANIEL JOSEPH) 12 September 2000 (2000-09-12) column 2, line 12 - line 40 column 3, line 48 - line 64 -----	59,60
A	HADDAD A ET AL: "TARGETED M CELL IMMUNIZATION FOR HIV-1 ENV DNA VACCINES" FASEB JOURNAL, FED. OF AMERICAN SOC. FOR EXPERIMENTAL BIOLOGY, BETHESDA, MD, US, vol. 14, no. 6, 20 April 2000 (2000-04-20), page A1204, XP000995418 ISSN: 0892-6638 abstract 185.7 -----	
E	WO 02/080852 A (O'MAHONY DANIEL J ;BRAYDEN DAVID J (IE); BYRNE DARAGH (IE); DIGITA) 17 October 2002 (2002-10-17) the whole document -----	1-58, 61-64

INTERNATIONAL SEARCH REPORT

International application No.
PCT/IB 02/03866

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.:
because they relate to subject matter not required to be searched by this Authority, namely:
Although partially claims 1-9, 12-21, 34-51, 61-64, and completely claims 10, 28-31, 33, are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. ☒ Claims Nos.: 1-58, 61-64 partially
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
see FURTHER INFORMATION sheet PCT/ISA/210
3. ☐ Claims Nos.:
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

see additional sheet

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☒ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
1-64

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
☐ No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Claims Nos.: 1-58, 61-64 partially

Present claims 1-58 and 61-64 relate to an extremely large number of possible compounds and the use thereof. In the present particular case, there is considered to be support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT for none of the compounds comprised in the claims. More particularly, there is a complete lack of evidence that the compounds recited in the claims provide any plausible solution to a technical problem possibly addressed by the application. Consequently, the claims are considered to so lack support, and the application to so lack disclosure, that a meaningful search over the whole of the claimed scope is impossible. As a result, the search has been carried out for those elements of the application which appear to be supported and disclosed, and for which a meaningful search was considered possible: in this particular case, a search was only considered possible on the inventive concept underlying the application, ie essentially that related to the correlation of altered expression of proteins in Peyer's patch cells / M cells with their use in enabling / facilitating oral delivery of drugs and antigens.

The applicant's attention is drawn to the fact that claims relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure. If the application proceeds into the regional phase before the EPO, the applicant is reminded that a search may be carried out during examination before the EPO (see EPO Guideline C-VI, 8.5), should the problems which led to the Article 17(2) declaration be overcome.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-64

Means to increase in PP cells of the intestine levels of proteins specific to PP cells, or decrease in PP cells of the intestine the levels of proteins non-specific to PP cells, via delivery of a nucleic acid encoding the PP cell specific protein / the protein per se, or delivery of an antisense / ribozyme / RNAi to the protein non-specific to PP cells, (eg claim 1 et seq., claim 15 and seq.); means to deliver a composition to a PP cell wherein said composition has a ligand that will specifically bind to a PP specific protein (eg claim 57); related subject matter

2. claim: 65

Promote enterocyte-M cell conversion via use of an antigen and a bacteria, probiotic yoghurt or bacterial component

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/IB 02/03866

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WO 02080852	A	17-10-2002	WO 02080852 A2	17-10-2002